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#### (FILE 'HOME' ENTERED AT 14:24:42 ON 06 AUG 2001) FILE 'HCAPLUS' ENTERED AT 14:24:48 ON 06 AUG 2001 L173 S ICT 45372 S IMMUNOASSAY? OR IMMUNOCHEMICAL ANALYSI? L2191 S IMMUNOCHROMATOG? (L) (ASSAY? OR IMMUNOASSAY?) L3114196 S L2 OR ASSAY? L4264 S L1 OR L3 L5101952 S ( H OR HAEMOPHILUS) L6 4047 S ( H OR HAEMOPHILUS) (2W) INFLUEN? L7 L81 S L7 AND L5 151595 S ANTIBOD? L9 181950 S ANTIGEN? L10 33182 S L9 (L) L10 L11L1294 S L11 AND L7 14 S L12 AND L4 L13 6647 S L4 (L) APP? L144 S L13 AND L14 L15 349 S (IMMUNOCHROMAT?)/AB L16 0 S L16 AND L7 L17 L18 399 S L3 OR L16 77 S L18 AND L9 AND L10 L19 99775 S CARBOHYDRAT? L20 43853 S POLYSACCHARID? L21 L22 139446 S L20 OR L21 4400 S L22 (L) L10 L23 5 S L23 AND L19 L24 29421 S STRIP# L25 0 S L12 AND L25 L26 L27 0 S L7 AND L4 AND L25 L28 2 S L7 AND L4 AND STRIP?/AB 7 SS L8 OR L15 OR L28 L29 FILE 'REGISTRY' ENTERED AT 14:34:33 ON 06 AUG 2001 E GOLD/CN L30 1 S E3 FILE 'HCAPLUS' ENTERED AT 14:34:40 ON 06 AUG 2001 L31 142553 S L30 OR GOLD OR AU L32 1159 S L31 AND L4 0 S L32 AND L7 L33 26 S L31 AND L3 L34 FILE 'STNGUIDE' ENTERED AT 14:35:57 ON 06 AUG 2001 FILE 'HCAPLUS' ENTERED AT 14:42:01 ON 06 AUG 2001 0 S L34 AND L7 L35 2 S L34 AND L23 L36 9 S L29 OR L36 L37

L37 ANSWER 1 OF 9 HCAPLUS COPYRIGHT 2001 ACS

=> d .ca 137 1-9



ACCESSION NUMBER:

2000:209941 HCAPLUS

DOCUMENT NUMBER:

132:233987

TITLE:

Process and materials for the rapid detection of

Streptococcus pneumoniae employing purified

antigen-specific antibodies

INVENTOR(S):

Moore, Norman James; Fent, Mary Kathleen; Koulchin,

Vladimir Andrei; Molokova, Elena Valentin

PATENT ASSIGNEE(S):

SOURCE:

Binax, Inc., USA PCT Int. Appl., 38 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent English

LANGUAGE:

FAMILY ACC. NUM. COUNT:

1

PATENT INFORMATION:

P.P.	TENT	NO.		KI	ND	DATE			A	PPLI	CATI	ON N	Ο.	DATE			
		<b>-</b> -							-								
WC	2000	0168	03	A	1	2000	0330		W	0 19	99-U	S215	05	1999	0920		
	W:	AT,	ΑU,	CA,	CH,	ĊN,	CZ,	DΕ,	DK,	ES,	FI,	GB,	HU,	IL,	IN,	JP,	KR,
		LU,	MX,	NO,	NZ,	PL,	PT,	RU,	SE,	SK,	UA,	zA					
	RW:	AT,	BE,	CH,	CY,	DE,	DK,	ES,	FI,	FR,	GB,	GR,	IE,	ΙŢ,	LU,	MC,	NL,
		PT,	SE			*											
ΑÜ	9961	513		Α	1	2000	0410		A	U 19	9.9-6	1513		1999	0920		
EF	1113	817		A	1 .	2001	0711		E	P 19	99-9	4830	5	1999	0920		
	R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	ΙT,	LI,	LU,	NL,	SE,	MC,	PT,
		ΙE,	FΙ														
PRIORIT	Y APP	LN.	INFO	.:					US 1	998-	1564	86	Α	1998	0918		
									US 1	999-	3971	10	Α	1999	0916		
								1	WO 1	999-1	US21	505	W	1999	0920		

A process is disclosed for obtaining a C-polysaccharide cell wall antigen AΒ contg. not more than about 10% protein from Streptococcus pneumoniae bacteria. The antigen thus obtained is conjugated to a spacer mol., and the free end of the latter is then conjugated to a chromatog. affinity column. The column is then utilized to purify raw antibodies to S. pneumoniae bacteria, thereby producing antigen-specific antibodies. A portion of such antibodies is conjugated to a labeling agent which displays a visible color change upon reaction of the antibodies with their

antigenic binding partner and embedded in a first zone of an immunochromatog. assay device. Another portion of such antibodies is bound to the reaction zone of the device which has a view window. When a liq. sample, such as patient urine, cerebrospinal fluid or blood is applied to the first zone, the conjugate of antibodies and labeling agent and the sample move along a flow strip of bibulous material to the reaction zone wherein, if the sample contains S. pneumoniae or its cell wall antigen, a sandwich is formed among the labeled conjugate, the antigen and the bound antibodies and a color change is obsd. The immunochromatog. assay thus performed is completed within about 15 min. This assay affords a basis for rapid and reliable diagnosis of various pathogenic states caused by S. pneumoniae including pneumonia, bronchitis,

otitis media, sinusitis, meningitis, and secondary disease states that commonly occur when primary pneumonic infection caused by this bacterium persists undiminished over a time period of 3-5 days.

IC ICM A61K039-385

ICS A61K039-085

CC 9-10 (Biochemical Methods)

ΙT Immunoassay (app., Immunochromatog. assay device; process and materials for rapid detection of Streptococcus pneumoniae employing purified antigen-specific antibodies) Polysaccharides, analysis ΙT Proteins, general, analysis RL: ANT (Analyte); THU (Therapeutic use); ANST (Analytical study); BIOL (Biological study); USES (Uses) (process and materials for rapid detection of Streptococcus pneumoniae employing purified antigen-specific antibodies) Carbohydrates, preparation IT RL: ARG (Analytical reagent use); PUR (Purification or recovery); ANST (Analytical study); PREP (Preparation); USES (Uses) (process and materials for rapid detection of Streptococcus pneumoniae employing purified antigen-specific antibodies) 302-01-2, Hydrazine, uses 7440-57-5, Gold, uses TΥ RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses) (process and materials for rapid detection of Streptococcus pneumoniae employing purified antigen-specific antibodies) REFERENCE COUNT: 11 (1) Bennett; The Journal of Immunology 1979, V122(6), REFERENCE(S): P2356 HCAPLUS (2) Fischer; European Journal of Biochemistry 1993, P851 HCAPLUS (4) Jennings; Biochemistry 1980, V19(20), P4712 **HCAPLUS** (5) Lees; US 5849301 A 1998 HCAPLUS (6) Marburg; US 4830852 A 1989 HCAPLUS ALL CITATIONS AVAILABLE IN THE RE FORMAT L37 ANSWER 2 OF 9 HCAPLUS COPYRIGHT 2001 ACS ACCESSION NUMBER: 1996:494700 HCAPLUS DOCUMENT NUMBER: 125:162734 Methods for detection of an analyte TITLE: Bogart, Gregory R.; Moddel, Garret R.; Maul, Diana INVENTOR(S): M.; Etter, Jeffrey B.; Crosby, Mark PATENT ASSIGNEE(S): Biostar, Inc., USA U.S., 71 pp. Cont.-in-part of U.S. Ser. No. SOURCE: 924343, abandoned. CODEN: USXXAM DOCUMENT TYPE: Patent English LANGUAGE: FAMILY ACC. NUM. COUNT: 14 PATENT INFORMATION: APPLICATION NO. DATE KIND DATE PATENT NO. ----------\_\_\_\_ А 19960730 US 1993-75952 19930610 US 5541057 19910320 A1 19921021 AU 1991-79004 AU 9179004 B2 19941020 AU 653940 A1 19930505 EP 539383 EP 1991-910056 19910320 B1 19960918 EP 539383 R: BE, CH, DE, ES, FR, GB, IT, LI, LU, NL, SE JP 05506936 T2 19931007 JP 1991-509344 19910320 ES 1991-910056 19910320 ES 2094224 T3 19970116

US 5639671

A 19970617

US 1995-412600

Page 3

19950328

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19950531
                                            US 1995-456040
     US 5629214
                       Α
                             19970513
                                            US 1995-455652
                                                              19950531
     US 5869272
                       Α
                             19990209
                                                              19980114
                                            JP 1998-5911
                       A2
                             19981027
     JP 10288616
                       B2
                             19990920
     JP 2951300
                                         US 1989-408291
                                                           B2 19890918
PRIORITY APPLN. INFO .:
                                                           B2 19920424
                                         US 1992-873097
                                                           B2 19920731
                                         US 1992-924343
                                                           A3 19900918
                                         JP 1990-513789
                                                           A 19910320
                                         EP 1991-910056
                                         WO 1991-US1781
                                                           A 19910320
                                         US 1992-923048
                                                           B2 19920731
                                         US 1993-75952
                                                           A3 19930610
                                         US 1993-76319
                                                           B1 19930610
     This invention relates to devices that produce a detectable attenuation
AΒ
of
     the spectral characteristic of light impinging on the devices by
thin-film
     phenomena. Interference phenomena are central to the devices and methods
     of the invention. The presence or amt. of an analyte of interest (e.g., rheumatoid factor, viral antigens, Streptococcus Group A antigen,
     allergens, HIV I or II, etc.) in a sample (e.g., blood, urine, spinal
     fluid, gastric wash, vaginal secretions, etc.) is found by using a
     substrate having an optically active surface exhibiting a first color in
     response to light impinging thereon and exhibiting a second color
     comprising a combination of wavelengths of light different from the first
     color or comprising an intensity of at least one wavelength of light
     different from the first color in response to the light when the analyte
     is present on the surface. Then the optically active surface is
contacted
     with a sample potentially comprising the analyte of interest under
     conditions in which the analyte can interact with the optically active
     surface to cause the optically active surface to exhibit the second color
     when the analyte is present. The devices permit detection of extremely
     small quantities of analyte in a sample, in amts. as low as 0.1 nM, 0.1
     ng/mL, 50 fg, or 2 .times. 103 organisms in a rapid assay that lasts only
     a few minutes.
     ICM C12Q001-70
IC
         G01N033-53; G01N033-543; G01N021-00
     ICS
NCL
     435005000
     9-1 (Biochemical Methods)
CC
     Section cross-reference(s): 3, 7, 15, 73
     optically active surface app biochem analysis; interference film optical
ST
     app biochem analysis; thin film analyzer body fluid; bacteria detection
     body fluid app; virus detection body fluid app; antigen
     detection body fluid app; antibody detection body fluid app
ΙT
     Bacteria
     Blood analysis
     Body fluid
     Cerebrospinal fluid
     Chlamydia
     Ellipsometers
     Escherichia coli
     Feces
     Films
     Immunoassay
     Infrared radiation
     Interference
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Latex
    Light
    Neisseria meningitidis
    Optical detectors
    Pericardium
    Peritoneum
    Pharynx
    Pleura
    Reflectometers
    Respiratory tract
    Saliva
    Sputum
    Stomach
    Streptococcus pneumoniae
    Ultraviolet radiation
    Urine analysis
        (app. and methods for anal. using thin-film phenomena)
    Haemophilus influenzae
IT
        (type b, app. and methods for anal. using thin-film phenomena)
    ANSWER 3 OF 9 HCAPLUS COPYRIGHT 2001 ACS
ACCESSION NUMBER: 1996:191556 HCAPLUS
DOCUMENT NUMBER:
                        124:337288
                        Devices and methods for detection of an analyte based
TITLE:
                       upon light interference
                       Sandstrom, Torbjorn; Stiblert, Lars; Maul; Diana M.
INVENTOR(S):
                       Biostar, Inc., USA
PATENT ASSIGNEE(S):
                        U.S., 69 pp., Cont.-in-part of U.S. Ser. No.
SOURCE:
923,268,
                        abandoned.
                        CODEN: USXXAM
                        Patent
DOCUMENT TYPE:
LANGUAGE:
                        English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:
                     KIND DATE
                                         APPLICATION NO. DATE
     PATENT NO.
                     _ _ _
                           _____
                                         -----
                                      US 1993-75128
US 1995-455493 19950531
19920731
                     A 19960227
    US 5494829
    US 5631171
                           19970520
                     Α
PRIORITY APPLN. INFO.:
                                       US 1993-75128
                                                          19930610
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AB Instrument configured and arranged to detect the presence or amt. of an analyte of interest on the substrate of an optical device is disclosed. The instrument has a source of linearly polarized, monochromatic light positioned at an angle other than Brewster's angle relative to the substrate; and an analyzer positioned at the same angle relative to the substrate at a location suitable for detecting reflected polarized light from the substrate; wherein the analyzer is configured and arranged to approx. maximize the change in intensity of the light reflected from the substrate that is transmitted through the analyzer when a change in mass occurs at the substrate relative to an unreacted surface.

IC ICM G01N033-543

ICS G01N

NCL 436518000

CC 9-1 (Biochemical Methods)

ST app light interference immunoassay; bacteria

antibody antigen detection app

ΙT Bacteria

Haemophilus influenzae

Neisseria meningitidis

Streptococcus pneumoniae

(devices and methods for detection of an analyte based upon light interference)

ΙT Immunoassay

(enzyme-linked immunosorbent assay, devices and methods for detection of an analyte based upon light interference)

L37 ANSWER 4 OF 9 HCAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER:

1996:184244 HCAPLUS

DOCUMENT NUMBER:

124:222848

TITLE:

Simultaneous detection, identification and differentiation of eubacterial taxa using a

hybridization assay

INVENTOR(S):

Jannes, Geert; Rossau, Rudi; Van Heuverswyn, Hugo

PATENT ASSIGNEE(S):

Innogenetics, N.V., Belg.

SOURCE:

PCT Int. Appl., 247 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

	PATENT NO.				KIND DATE			APPLICATION NO. DATE										
	WO 9600298			A1 19960104				WO 1995-EP2452 19950623										
															DK,			FI,
															LT,			
															SG,			
			TM,															
		RW:	ΚE,	MW,	SD,	SZ,	UG,	ΑT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	ΙE,	IT,
			LU,	MC,	NL,	PT,	SE,	BF,	ВJ,	CF,	CG,	CI,	CM,	GΑ,	GN,	ML,	MR,	ΝE,
			SN,	TD,	TG													
	CA 2193101				A	A	1996	0104		CA 1995-2193101 19950623								
	AU 9529246			A1 19960119				AU 1995-29246 19950623 EP 1995-924923 19950623										
	EΡ																	
		R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	ΙE,	IT,	LI,	LU,	MC,	NL,	PT,
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	BR	9508	101		A	_	1997	1230		В.	R 19	95-8. 05-5	7000 TOT	,	1995	0623		
	JP	1050	19/6		T2 19980224				JP 1995-502804 19950623 EP 2001-200037 19950623									
	EP	10888	899		A2 20010404 A3 20010502			0404		Ľ.	P 20	01-21	0003	1	1990	0623		
	EР									CD	CD	TT	тт	T II	NIT	C F	мс	рπ
		к:	-				DK,	LS,	rr,	GD,	GR,	11,	ידר,	що,	NL,	JE,	ric,	ΕΙ,
	EВ	1098			LT,		2001	n5n9		E.	P 20	01-2	nnn4	5	1995	0623		
	EP														NL,		MC.	PΤ,
		κ.			LT,			ц,	111,	OD,	OI.,	,	,		112,	02,	,	,
	EP	1098	007	01,	Δ1,	1	2001	0509		E	P 20	01-2	0004	6	1995	0623		
	בים														NL,		MC,	PT,
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	EP 1091004			A2 20010411			EP 2001-200042 19950624											
		1091																
		R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,
ΙE																		

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US 1996-765332
                                                               19961223
                             20000215
     US 6025132
                        Α
                             19990902
                                            AU 1999-35035
                                                               19990615
                        Α1
     AU 9935035
                             20000907
                        B2
     AU 723742
                                          EP 1994-870106
                                                               19940624
                                                            Α
PRIORITY APPLN. INFO.:
                                                               19950407
                                          EP 1995-870032
                                                            Α
                                          AU 1995-29246
                                                            A3 19950623
                                          EP 1995-924923
                                                            A3 19950623
                                          WO 1995-EP2452
                                                            W 19950623
     A method is presented for detection and identification of at least one
AΒ
     microorganism, or for the simultaneous detection of several
microorganisms
     in a sample. The polynucleic acids present in the sample may need to be
     released, isolated, or concd., and if necessary, the 16S-23S rRNA spacer
     region, or a part of it, is amplified with at least one suitable primer
     pair. The polynucleic acids are hybridized with at least one and
     preferably more than one of the 112 spacer probes or their equiv., under
     the appropriate hybridization and wash conditions, and/or with a
     taxon-specific probe derived from any of 103 spacer sequences under the same hybridization and wash conditions. The hybrids formed with each of
     the probes used under appropriate hybridization and wash conditions are
     detected, the microorganism(s) present in the sample identified from the
     differential hybridization signals obtained.
     ICM C12Q001-68
IC
CC
     3-1 (Biochemical Genetics)
     Section cross-reference(s): 10
IT
     Acinetobacter
     Acinetobacter baumannii
     Bordetella pertussis
     Brucella
     Campylobacter
     Cerebrospinal fluid
     Chlamydia
     Chlamydia psittaci
     Chlamydia trachomatis
     Digestive tract
     Food analysis
     Genitourinary tract
     Haemophilus ducreyi
     Haemophilus influenzae
     Listeria
     Listeria monocytogenes
     Microorganism
     Moraxella catarrhalis
     Mycobacterium
     Mycobacterium avium
     Mycobacterium celatum
     Mycobacterium chelonae
     Mycobacterium fortuitum
     Mycobacterium genavense
     Mycobacterium gordonae
     Mycobacterium haemophilum
     Mycobacterium intracellulare
     Mycobacterium kansasii
     Mycobacterium malmoense
     Mycobacterium marinum
     Mycobacterium paratuberculosis
     Mycobacterium scrofulaceum
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Mycobacterium simiae Mycobacterium tuberculosis Mycobacterium ulcerans Mycobacterium xenopi Mycoplasma Mycoplasma genitalium Mycoplasma pneumoniae Neisseria gonorrhoeae Neisseria meningitidis Pseudomonas Pseudomonas aeruginosa Respiratory tract Salmonella Staphylococcus Staphylococcus aureus Staphylococcus epidermidis Streptococcus Streptococcus agalactiae Streptococcus pneumoniae Yersinia enterocolitica (simultaneous detection, identification and differentiation of eubacterial taxa using a hybridization assay based on the 16 S-23 S rRNA spacer region) IT174696-37-8 RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses) (probe CHPS-ICT 1; simultaneous detection, identification and differentiation of eubacterial taxa using a hybridization assay based on the 16 S-23 S rRNA spacer region) IT 174696-34-5 RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses) (probe CHTR-ICT 1; simultaneous detection, identification and differentiation of eubacterial taxa using a hybridization assay based on the 16 S-23 S rRNA spacer region) ΤТ 174696-35-6 .RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses) (probe CHTR-ICT 2; simultaneous detection, identification and differentiation of eubacterial taxa using a hybridization assay based on the 16 S-23 S rRNA spacer region) IT 174696-36-7 RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses) (probe CHTR-ICT 3; simultaneous detection, identification and differentiation of eubacterial taxa using a hybridization assay based on the 16 S-23 S rRNA spacer region) L37 ANSWER 5 OF 9 HCAPLUS COPYRIGHT 2001 ACS 1995:774831 HCAPLUS ACCESSION NUMBER: 123:164647 DOCUMENT NUMBER: Interrupted-flow assay device TITLE: Chandler, Howard M. INVENTOR (S): Smithkline Diagnostics, USA PATENT ASSIGNEE(S):

PCT Int. Appl., 76 pp.

CODEN: PIXXD2

Patent English

SOURCE:

LANGUAGE:

DOCUMENT TYPE:

FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

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KIND DATE
     PATENT NO.
                                          APPLICATION NO. DATE
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                                          _____
                     A1 19950615
                                          WO 1994-US14004 19941206
    WO 9516208
            AM, AT, AU, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, ES, FI, GB,
            GE, HU, JP, KE, KG, KP, KR, KZ, LK, LT, LU, LV, MD, MG, MN, MW,
            NL, NO, NZ, PL, PT, RO, RU, SD, SE, SI, SK, TJ, TT, UA, US, UZ,
VN
        RW: KE, MW, SD, SZ, AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU,
            MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN,
            TD, TG
                           19951121
                                          US 1993-163341
                                                           19931207
    US 5468648
                                          AU 1995-13008
                                                           19941206
    AU 9513008
                      Α1
                           19950627
    AU 684585
                      B2
                           19971218
                                          EP 1995-904245
                      Α1
                           19960925
                                                           19941206
    EP 733211
    EP 733211
                      В1
                          19980513
        R: BE, CH, DE, ES, FR, GB, IT, LI, NL, SE
                                          JP 1994-516281
                                                           19941206
                      T2 19970617
     JP 09506177
                                       US 1993-163341 A 19931207
PRIORITY APPLN. INFO.:
                                       US 1991-706639
                                                       A2 19910529
                                       US 1992-888831
                                                       B2 19920527
                                       US 1993-40430
                                                        A2 19930331
                                       WO 1994-US14004 W 19941206
    The present invention provides chromatog. assay devices that can perform
AΒ
    multiple assays simultaneously in the same test strip, as well as methods
    for their use. One of the assays can be an immunol. assay to detect an
    antigen, such as human chorionic gonadotropin, while another assay can be
    a serol. assay to detect an antibody, such as antirubella antibody. An
    assay device according to the present invention can comprise: (1) a first
    opposable component including at least one chromatog. medium having a
    specific binding partner to the first analyte and a specific binding
    partner to the second analyte immobilized thereto in sep., discrete,
    non-overlapping zones; and (2) a second opposable component including an
    absorber. The first and second opposable components are configured such
    that bringing the first and second opposable components into opposition
    causes the absorber to come into operable contact with at least one
    chromatog. medium so that the zone contg. the specific binding partner to
    the first analyte is functionally divided from the zone contg. the
    specific binding partner to the second analyte so that both analytes can
    be detected.
IC
    ICM G01N033-558
    ICS G01N033-543
    9-1 (Biochemical Methods)
CC
    Section cross-reference(s): 14, 15
ST
    interrupted flow assay app biochem analysis; antibody antigen
    detn interrupted flow app; immunochromatog assay
    antibody antigen detn; chromatog immunoassay app
    Carbohydrates and Sugars, analysis
ΙT
    Glycoproteins, analysis
    Haptens
    Hemoglobins
    Mucoproteins
    Proteins, analysis
    RL: ANT (Analyte); ANST (Analytical study)
        (interrupted-flow assay app. for antibody and antigen detn.)
     7440-57-5, Gold, uses
    RL: ARG (Analytical reagent use); ANST (Analytical study); USES (Uses)
        (interrupted-flow assay app. for antibody and antigen detn.)
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L37 ANSWER 6 OF 9 HCAPLUS COPYRIGHT 2001 ACS ACCESSION NUMBER: 1993:467251 HCAPLUS

DOCUMENT NUMBER: 119:67251

TITLE: Highly sensitive optical immunoassay using

enzyme-labeled reagents

INVENTOR(S): Maul, Diana M.; Crider, Debbie G.; Bilodeau, Robert

J.; Bogart, Gregory R. Biostar, Inc., USA

PATENT ASSIGNEE(S): Biostar, Inc., USA SOURCE: Can. Pat. Appl., 37 pp.

CODEN: CPXXEB

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 14

PATENT INFORMATION:

PA'	TENT NO.	KIND	DATE	APPLICATION NO. DATE
CA	2078897			CA 1992-2078897 19920923
				AU 1991-79004 19910320
	653940			
	539383		19930505	EP 1991-910056 19910320
	539383			*
				IT, LI, LU, NL, SE
JP	05506936	T2		JP 1991-509344 19910320
ES	2094224	Т3	19970116	ES 1991-910056 19910320
				EP 1991-308968 19911001
EP	546222	В1	19970910	
				FR, GB, GR, IT, LI, LU, NL, SE
AT	158080	E	19970915	AT 1991-308968 19911001 ES 1991-308968 19911001
ES	2109258	Т3	19980116	ES 1991-308968 19911001
US	5955377	Α	19990921	US 1995-403565 19950417
PRIORIT	Y APPLN. INFO	.:		EP 1991-308968 A 19911001
				US 1991-653052 A2 19910211
				EP 1991-910056 A 19910320
				WO 1991-US1781 A 19910320
				US 1992-923090 B2 19920731
				US 1993-75693 B1 19930610

AB A thin-film optical immunoassay device is disclosed which comprises a solid support substrate having an upper and lower surface, an unlabeled ligand antibody layer bound to the substrate, .gtoreq.1 layer comprising an immobilized enzyme conjugate, complexed with an analyte of interest

and
capable of further interacting with an enzyme-reactive delivery substance
to form an insol. reaction product. The enzyme conjugate layer and the
unlabeled antibody layer have a measurable increased mass change, the

capable of pptn. by a pptg. agent applied as a substrate thereover. Also provided are a corresponding process for detection of an analyte in a medium and a diagnostic test kit for a thin-film optical immunoassay.

invention overcomes the limitations imposed by the prior art use of particulate reagent enhancers. By the use of enzyme-antibody conjugates in place of latex-reagent particles, even more highly sensitive optical thin-film assays can be obtained, particularly with selected substrates for the enzyme which provide insol. pptd. products. The invention

relates

to the use of such enzyme-labeled antibody methods in thin-film assays for the detection of low levels of the polysaccharide antigens derived from the group of bacteria commonly responsible for bacterial infections in man, e.g. streptococcus. Thus, a conjugate of peroxidase with Igs from antisera against Neisseria meningitidis (A, C, Y, W135) was dild. in casein-contg. buffer and mixed with an equal vol. of a diln. of a cell-free filtrate from a culture of Neisseria meningitidis. The mixt. was pipeted onto the surface of a silicon wafer already coated with layers of silicon nitride, t-polymer siloxane, and purified Ig from the same anti-N. meningitidis antiserum. TMBlue was used as pptg. substrate; the ppt. was read by eye and by ellipsometer to confirm the presence of N. meningitidis. Visually, a 1:20,000 diln. of antigen was clearly resolved from the neg.; a com. kit's sensitivity cut-off is a 1+ at a 1:4000 diln. of the same antigen prepn. IC ICM G01N033-569 9-1 (Biochemical Methods) CCthin film optical immunoassay; enzyme antibody conjugate optical STimmunoassay; Neisseria thin film optical immunoassay; Streptococcus thin film optical immunoassay ΙT Bacteria Streptococcus pneumoniae (analyte derived from, detn. of, thin-film optical immunoassay device with enzyme-antibody conjugate for, increased sensitivity in relation to) IT Antigens Polysaccharides, analysis RL: ANST (Analytical study) (bacteria-derived, detn. of, thin-film optical immunoassay device with enzyme-antibody conjugate for, increased sensitivity in relation to) ΙT Ellipsometry (for thin-film optical immunoassay device with enzyme-antibody conjugate, increased sensitivity in relation to) IT Immobilization, biochemical (of unlabeled antibody and enzyme conjugate, for optical thin-film immunoassay device, increased sensitivity in relation to) ΙT Films (thin, optical immunoassay device using, with antibody-enzyme conjugate, increased sensitivity in relation to) TT Immunoassay (app., thin-film optical, with antibody-enzyme conjugate, increased sensitivity in relation to) TT Immunoglobulins RL: ANST (Analytical study) (conjugates, anti-Neisseria meningitidis, with peroxidase, for optical thin-film immunoassay of N. meningitidis antigens) IT Enzymes RL: ANST (Analytical study) (conjugates, with antibodies, for thin-film optical immunoassay device, increased sensitivity in relation to) IT Antibodies RL: ANST (Analytical study) (conjugates, with enzymes, for thin-film optical immunoassay device, increased sensitivity in relation to) IT Neisseria meningitidis

```
Streptococcus
        (group A, analyte derived from, detn. of, thin-film optical
      immunoassay device with enzyme-antibody conjugate for,
        increased sensitivity in relation to)
ΙT
     Neisseria meningitidis
     Streptococcus
        (group B, analyte derived from, detn. of, thin-film optical
      immunoassay device with enzyme-antibody conjugate for,
        increased sensitivity in relation to)
TΤ
     Neisseria meningitidis
        (group C, analyte derived from, detn. of, thin-film optical
      immunoassay device with enzyme-antibody conjugate for,
        increased sensitivity in relation to)
ΤΤ
     Neisseria meningitidis
        (group W-135, analyte derived from, detn. of, thin-film optical
      immunoassay device with enzyme-antibody conjugate for,
        increased sensitivity in relation to)
     Neisseria meningitidis
ΙT
        (group Y, analyte derived from, detn. of, thin-film optical
      immunoassay device with enzyme-antibody conjugate for,
        increased sensitivity in relation to)
ΙT
     Antibodies
     RL: ANST (Analytical study)
        (monoclonal, to bacteria, detn. of, optical thin-film
      immunoassay device for)
     Haemophilus influenzae
IT
        (type b, analyte derived from, detn. of, thin-film optical
      immunoassay device with enzyme-antibody conjugate for,
        increased sensitivity in relation to)
     54827-17-7, 3,3',5,5'-Tetramethylbenzidine
ΤT
     RL: ANST (Analytical study)
        (as pptg. agent, in thin-film optical immunoassay with
        antibody-enzyme conjugate)
     9003-99-0D, Peroxidase, anti-bacterial antibody conjugates
IT
     RL: ANST (Analytical study)
        (for thin-film optical immunoassay device)
     12033-89-5, Silicon nitride, biological studies
IT
     RL: BIOL (Biological study)
        (silicon wafer coated with, optical thin-film immunoassay in
        relation to)
     7440-22-4, Silver, biological studies
ΤТ
     RL: BIOL (Biological study)
        (silicon wafer colored with, optical thin-film immunoassay in
        relation to)
     7440-21-3, Silicon, biological studies
ΙT
     RL: BIOL (Biological study)
        (wafer, silicon nitride-coated or silver-colored, optical thin-film
      immunoassay in relation to)
L37 ANSWER 7 OF 9 HCAPLUS COPYRIGHT 2001 ACS
                         1987:64041 HCAPLUS
ACCESSION NUMBER:
DOCUMENT NUMBER:
                         106:64041
                         Solid-phase system and apparatus for use in
TITLE:
                         ligand-receptor assays, particularly
                       immunoassays
                         Rubenstein, Albert Samuel
INVENTOR(S):
PATENT ASSIGNEE(S):
                         Hybritech, Inc., USA
```

SOURCE:

Eur. Pat. Appl., 30 pp.

CODEN: EPXXDW

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

P.F	ATENT NO.		KIND	DATE		APPLICATION NO.	DATE
E F	200381		A1	19861105		EP 1986-302521	19860404
	200381			19930127			
	R: AT,	BE,				I, LU, NL, SE	•
	1272127					CA 1986-505788	
F]	8601460 92257 92257		A	19861005		FI 1986-1460	19860404
F]	92257		В	19940630			
F	92257		С	19941010			
DF	8601538		A	19861005		DK 1986-1538	19860404
	( 171928		В1	19970811			
NC	8601317		A	19861006		NO 1986-1317	19860404
	170825		В	19920831			
NC	170825		С	19921209			
ΑU	8655657		A1	19861016		AU·1986-55657	19860404
ΑU	0 605101		B2	19910110			
JI	61292059		A2	19861222		JP 1986-78064	19860404
JI	07113636		B4	19951206			
ES	5 553724		A1	19890201		ES 1986-553724	19860404
ES	5 553724			19890331			
E	437287			19910717		EP 1991-103886	19860404
E	437287		A3	19910724			
E	437287		B1	19960717			
	R: AT,	BE,	CH, DE	, FR, GB,	IT, L	I, LU, NL, SE	
ΑT	85129		E	19930215		AT 1986-302521 AT 1991-103886	19860404
A	140539		E	19960815		AT 1991-103886	19860404
	5879881			19990309		US 1993-162791	
PRIORI	Y APPLN.	INFO	.:		US	1985-720036	19850404
					US	1986-847799	19860403
					EP	1986-302521 1987-3496 1990-579087 1992-879693	19860404
					US	1987-3496	198/0115
					US	1990-579087	19900907
א פא	ealid-pha	9 9	vetem f	or use in	title	assavs (e.g., im	munoassavs.

A solid-phase system for use in title assays (e.g., immunoassays, nucleic acid probe assays) comprises a porous matrix in which microspheres, bound to a receptor capable of capturing a target ligand, are entrapped. Microspheres with different bound receptors may be entrapped at different sites in the matrix enabling the detn. of .gtoreq.2 analytes in a sample at the same time. Also, internal controls may be incorporated by having zones of microspheres without receptors. For the detection of human chorionic gonadotropin (HCG) in urine, polystyrene microspheres were activated with monoclonal antibodies to HCG and then entrapped in Whatman GF/F glass fiber filters incorporated in an immunoconcn. app. Urine, followed by alk. phosphatase-labeled monoclonal antibodies to HCG, were applied to the filter and drawn through. The filter was washed and indoxyl phosphate was added. A dark blue color developed in the discrete zone of activated microspheres, indicating the presence of HCG.

IC ICM G01N033-549

G01N033-545; G01N033-551; G01N033-577; G01N033-564; G01N033-569; G01N033-74; G01N033-70

```
9-10 (Biochemical Methods)
CC
     Section cross-reference(s): 2, 15
     solid phase ligand receptor assay; chorionic gonadotropin solid
ST
     phase EIA; microsphere ligand receptor assay
TΤ
     Bacteria
     Candida albicans
     Chlamydia trachomatis
     Fungi
     Haemophilus influenzae
     Neisseria gonorrhoeae
     Parasite
     Trichomonas vaginalis
     Virus, animal
        (detn. of, matrix-entrapped receptor-bound microspheres in
      immunoassay for)
ΙT
     Receptors
     RL: ANST (Analytical study)
        (matrix-entrapped microsphere-bound, for ligands, detn. of, by
        solid-phase assay)
IT
     Allergens
     Antigens
     RL: ANST (Analytical study)
        (matrix-entrapped microsphere-bound, in solid-phase immunoassay
TΤ
     Latex
        (matrix-entrapped microspheres, receptor-bound, in solid-phase
      immunoassay)
ΙT
     Blood analysis
     Urine analysis
        (matrix-entrapped receptor-bound microspheres in solid-phase
      immunoassay for)
ΙT
     Ceramic materials and wares
     Filtering materials
     Glass fibers, uses and miscellaneous
     Polyamide fibers, uses and miscellaneous
     RL: USES (Uses)
        (receptor-bound microspheres entrapped in, for solid-phase
      immunoassay)
ΙT
     Ligands
     RL: ANST (Analytical study)
        (receptor-bound microspheres for, porous matrix-entrapped, in
        solid-phase assay)
ΙT
     Antibodies
     RL: PROC (Process)
        (to rubella virus, detn. of, with matrix-entrapped receptor-bound
        microspheres in solid-phase immunoassay)
IT
     Immunoglobulins
     RL: ANT (Analyte); ANST (Analytical study)
        (E, detn. of, matrix-entrapped allergen-bound microspheres in
      immunoassay for)
ΙT
     Virus, animal
        (adeno-, detn. of, matrix-entrapped receptor-bound microspheres in
      immunoassay for)
IT
     Antigens
     RL: ANT (Analyte); ANST (Analytical study)
        (carcinoembryonic, detn. of, matrix-entrapped receptor-bound
        microspheres in immunoassay for)
```

```
ΙT
     Virus, animal
        (cytomegalo-, detn. of, matrix-entrapped receptor-bound microspheres
in
      immunoassay for)
ΙT
     Immunochemical analysis
        (enzyme immunoassay, solid-phase, matrix-entrapped
      antigen- or antibody-bound microspheres in)
ΙT
     Streptococcus
        (group A, detn. of, matrix-entrapped receptor-bound microspheres in
      immunoassay for)
ΙT
     Streptococcus
        (group B, detn. of, matrix-entrapped receptor-bound microspheres in
      immunoassay for)
ΙΤ
     Virus, animal
        (hepatitis, detn. of, matrix-entrapped receptor-bound microspheres in
      immunoassay for)
ΙΤ
     Virus, animal
        (hepatitis A, detn. of, matrix-entrapped receptor-bound microspheres
in
      immunoassay for)
ΙΤ
     Virus, animal
        (hepatitis B, detn. of, matrix-entrapped receptor-bound microspheres
in
      immunoassay for)
ΙT
     Virus, animal
        (hepatitis, non-A, non-B, detn. of, matrix-entrapped receptor-bound
        microspheres in immunoassay for)
ΙT
     Virus, animal
        (herpes, detn. of, matrix-entrapped receptor-bound microspheres in
      immunoassay for)
     Virus, animal
IT
        (human T-cell leukemia, detn. of, matrix-entrapped receptor-bound
        microspheres in immunoassay for)
     Immunochemical analysis
IT
        (immunoassay, solid-phase, matrix-entrapped antigen
        - or antibody-bound microspheres in)
     Virus, animal
ΙT
        (influenza, detn. of, matrix-entrapped receptor-bound microspheres in
      immunoassay for)
TΤ
     Antigens
     RL: ANT (Analyte); ANST (Analytical study)
        (prostate-specific, detn. of, matrix-entrapped receptor-bound
        microspheres in immunoassay for)
ΙT
     Virus, animal
        (respiratory syncytial, detn. of, matrix-entrapped receptor-bound
        microspheres in immunoassay for)
     Virus, animal
ΙT
        (rota-, detn. of, matrix-entrapped receptor-bound microspheres in
      immunoassay for)
ΙT
     Virus, animal
        (rubella, detn. of, matrix-entrapped receptor-bound microspheres in
      immunoassay for)
ΙT
     Fetoproteins
     RL: ANT (Analyte); ANST (Analytical study)
        (.alpha.-, detn. of, matrix-entrapped receptor-bound microspheres in
      immunoassay for)
                9002-61-3, Choriogonadotropin
                                                  9002-67-9
IT
     9001-77-8
                                                                        Page 15
```

```
RL: ANT (Analyte); ANST (Analytical study)
        (detn. of, matrix-entrapped receptor-bound microspheres in
      immunoassay for)
ΙT
     9001-15-4
     RL: ANST (Analytical study)
        (isoenzymes, detn. of, matrix-entrapped receptor-bound microspheres in
      immunoassay for)
                               9003-07-0, Polypropylene
                                                          9003-53-6,
     9002-88-4, Polyethylene
ΙT
     Polystyrene
     RL: ANST (Analytical study)
        (matrix-entrapped microspheres, receptor-bound, in solid-phase
      immunoassay)
L37 ANSWER 8 OF 9 . HCAPLUS COPYRIGHT 2001 ACS
                         1985:75210 HCAPLUS
ACCESSION NUMBER:
                         102:75210
DOCUMENT NUMBER:
                         Immunoblot method for identifying surface components,
TITLE:
                         determining their cross-reactivity, and investigating
                         cell topology: results with Haemophilus
                       influenzae type b
                         Loeb, Marilyn R.
AUTHOR (S):
                         Med. Cent., Univ. Rochester, Rochester, NY, 14642,
CORPORATE SOURCE:
USA
                         Anal. Biochem. (1984), 143(1), 196-204
SOURCE:
                         CODEN: ANBCA2; ISSN: 0003-2697
                         Journal
DOCUMENT TYPE:
                         English
LANGUAGE:
    An outer membrane prepn. derived from encapsulated (type b) H. influenzae
    was resolved by SDS-polyacrylamide gel electrophoresis, and the sepd.
     components were then transferred electrophoretically to nitrocellulose.
    The nitrocellulose was cut into vertical strips, which were then
     each incubated with rabbit antiserum to the whole bacterium or with the
     same antiserum after absorption with any of the following: the same
     of H. influenzae, a capsule-deficient mutant of that strain, other
strains
    of H. influenzae, or other bacteria. The strips were then
     incubated with 125I-protein A, and the bound antibodies were detected by
     autoradiog. The autoradiograph of the strip exposed to
    unabsorbed antisera revealed the identity of those individual outer
    membrane components that bound antibodies. A comparison of the intensity
    of the various bands on this strip with those on the
     strips exposed to adsorbed antisera was then used to identify: (1)
     surface-exposed components, (2) those components occluded by capsule, and
     (3) cross-reactivity of exposed components. This method should be
     applicable to other cells and subcellular particles. Its major
     disadvantage is that it can provide false neg. results.
CC
     9-10 (Biochemical Methods)
     Section cross-reference(s): 10
ΙT
    Lipopolysaccharides
     Proteins
     RL: ANT (Analyte); ANST (Analytical study)
        (detection of, of Haemophilus influenzae surface
        membrane by gel electrophoresis and immunoblot)
     Immunochemical analysis
        (gel electrophoresis combined with, of lipopolysaccharides and
proteins
                                                                        Page 16
```

of Haemophilus influenzae surface membrane) ΙT Haemophilus influenzae (lipopolysaccharides and proteins in surface membranes of, by gel electrophoresis and immunoblot) Electrophoresis and Ionophoresis IT (gel, immunoblot combined with, of lipopolysaccharides and proteins of Haemophilus influenzae surface membrane) ΙT Cell wall (outer membrane, lipopolysaccharides in proteins detection in, of Haemophilus influenzae surface membrane by gel electrophoresis and immunoblotting) L37 ANSWER 9 OF 9 HCAPLUS COPYRIGHT 2001 ACS 1984:625331 HCAPLUS ACCESSION NUMBER: DOCUMENT NUMBER: 101:225331 Pyridinium 2-azo-p-dimethylaniline chromophore, a TITLE: chromogenic reagent for .beta.-lactamase testing compared to nitrocefin Barlam, T.; Neu, H. C. AUTHOR(S): Coll. Phys. Surg., Columbia Univ., New York, NY, CORPORATE SOURCE: 10032, USA Eur. J. Clin. Microbiol. (1984), 3(3), 185-9 SOURCE: CODEN: EJCMDM; ISSN: 0722-2211 DOCUMENT TYPE: Journal English LANGUAGE: Pyridinium-2-azo-p-dimethylaniline chromophore was evaluated as a test AB tube, filter paper, and spectrometric assay for detection of .beta.-lactamases from gram-pos. and gram-neg. organisms. Although useful for detection of TEM .beta.-lactamases in Haemophilus influenzae and Neisseria gonorrhoeae, it was a poor agent for detecting TEM, OXA, and PSE enzymes in Enterobacteriaceae. It also proved poor for detecting cephalosporinases in Pseudomonas aeruginosa and Enterobacteriaceae, and penicillinases in Staphylococcus aureus when compared to nitrocefin. As а spectrometric substrate it was equiv. to nitrocefin and cephaloridine with various .beta.-lactamases, but it was not useful in the above assays with test tubes or filler paper strips. CC 7-1 (Enzymes) Section cross-reference(s): 9, 10 IT Bacteria Haemophilus influenzae Neisseria gonorrhoeae (.beta.-lactamase of, detn. of, azodimethylaniline chromophore as substrate for) IT9001-74-5 9012-26-4 RL: ANT (Analyte); ANST (Analytical study) (detn. of, azodimethylaniline chromophore as substrate for, of bacteria, spectrometric assay in relation to)

## => fil wpids

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MOST RECENT DERWENT UPDATE 200143 <200143/DW>
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(FILE 'WPIDS' ENTERED AT 14:44:55 ON 06 AUG 2001)
                DEL HIS Y
            110 S IMMUNOCHROMATOG?
L1
           33 S ICT
L2
           140 S L1 OR L2
L3
        24802 S ANTIGEN?
L4
L5
          39493 S ANTIBOD?
             42 S L3 AND L4 AND L5
L6
           1425 S H INFLUENZ? OR HAEMOPHI?
L7
              0 S L6 AND L7
L8
              0 S L7 AND L3
L9
L10
            138 S L4 AND L5 AND L7
          27593 S ASSAY? OR IMMUNOASSAY? OR IMMUNOCHEM?
L11
             38 S L10 AND L11
L12
L13
        1562337 S APP## OR APPARAT? OR STRIP#
              1 S L12 AND L13
L14
L15
          74565 S AU OR GOLD
L16
              2 S L12 AND L15
            535 S (CARBOHYDRAT## OR ?SACCHARIDE?) (4A) L4
L17
              6 S L12 AND L17
L18
              7 S L14 OR L16 OR L18
L19
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FILE 'WPIDS' ENTERED AT 14:50:46 ON 06 AUG 2001

=> d .wp 1-7

L19 ANSWER 1 OF 7 WPIDS COPYRIGHT 2001 DERWENT INFORMATION LTD AN 2001-328612 [34] WPIDS DNC C2001-100777

```
Determining the dose response of a human to a bacterial polysaccharide
ΤI
    conjugate vaccine comprises measuring an immune response.
DC
    LAFERRIERE, C A J; POOLMAN, J; SLAOUI, M M
IN
     (SMIK) SMITHKLINE BEECHAM BIOLOGICALS
PΑ
CYC
    WO 2001030390 A2 20010503 (200134)* EN
                                              .35p
PΙ
        RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ
            NL OA PT SD SE SL SZ TZ UG ZW
        W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CR CU CZ DE DK DM
            DZ EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC
            LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE
            SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW
    WO 2001030390 A2 WO 2000-EP10733 20001027
                      19991028
PRAI GB 1999-25559
    WO 200130390 A UPAB: 20010620
AB
    NOVELTY - A method (I) of determining the dose response of a human to a
    polysaccharide conjugate vaccine, is new,
          DETAILED DESCRIPTION - A method (I) of determining the dose response
    of a human to a polysaccharide conjugate vaccine comprising an
immunogenic
    carrier protein and a bacterial polysaccharide. The method comprises:
          (a) administering a dose of the conjugate vaccine to an infant
    animal; and
          (b) determining the immune response of the animal to the bacterial
     polysaccharides as a measure of the immune response of a human.
          INDEPENDENT CLAIMS are included for the following:
          (1) The use of animals in (I); and
          (2) A combination vaccine (II) comprising 2 or more pneumococcal
    capsular polysaccharide conjugate antigens at an
    optimal concentration for inducing an optimal anti-polysaccharide
    antibody response when administered to a human;
          ACTIVITY - Antibacterial.
          MECHANISM OF ACTION - Vaccine.
          No supporting data is given.
          USE - (II) can be used to treat pneumococcal disease in a human host
    and for the manufacture of medicaments for treatment (both claimed).
          The method (I) is used for testing a vaccine response in an animal
    model to obtain information on the response to humans to the same vaccine
     antigen (claimed). The animals may be used in potency tests for
     the lot release of batches of vaccines to ensure that a related response
     in humans would be acceptable, and in pre-clinical studies to evaluate
the
    efficacy of new formulations of conjugate without initially having to
    conduct human trials. The animal model is used to develop a combination
    vaccine comprising 2 or more pneumococcal capsular polysaccharide
    conjugate antigens at an optimal concentration of inducing an
     optimal anti-polysaccharide antibody response when administered
          ADVANTAGE - The animal model is highly predictive of the human
     response to the antigen.
     Dwg.0/12
    ANSWER 2 OF 7 WPIDS COPYRIGHT 2001
                                           DERWENT INFORMATION LTD
     2000-442428 [38]
                        WPIDS
AN
                        DNC C2000-134585
    N2000-330108
DNN
     New biosensors for detection of analytes, especially yeast, fungi,
TТ
```

bacteria, or virus, comprises an antibody-binding protein layer

printed in a pattern onto a polymer film and an antibody attached to it. A18 A28 A96 B04 D16 J04 M13 S03 EVERHART, D S; KAYLOR, R M; MCGRATH, K (KIMB) KIMBERLY-CLARK WORLDWIDE INC CYC WO 2000036416 A1 20000622 (200038)\* EN 34p PΙ RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL OA PT SD SE SL SZ TZ UG ZW W: AE AL AM AT AU AZ BA BB BG BR BY CA CH CN CR CU CZ DE DK DM EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG UZ VN YU ZA ZW AU 2000017431 A 20000703 (200046) ADT WO 2000036416 A1 WO 1999-US27727 19991122; AU 2000017431 A AU 2000-17431 19991122 AU 2000017431 A Based on WO 200036416 PRAI US 1998-213713 19981217 WO 200036416 A UPAB: 20000811 NOVELTY - A biosensor comprising a polymer film, and an antibody -binding protein (ABP) layer printed in a pattern onto the polymer film, where the ABP layer has an antibody on it that is specific for an analyte, is new. DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following: (1) making a biosensor comprising printing a pattern of ABP layer with a subsequent layer of antibody on the polymer film; (2) detecting an analyte in a medium, comprising contacting the medium with a biosensing device of the novelty, transmitting light through the polymer film, and detecting the presence of the analyte bound to the antibody, by detecting a pattern form by diffraction of the light; and (3) a biosensor comprising a polymer film, and an ABP layer printed in a pattern onto the polymer film where the ABP layer is capable of acting as a receptor for an analyte. USE - The biosensors can be used for the detection of analytes such as bacteria, yeast, fungus, virus, rheumatoid factor, IgG, IgM, IgA and IgE antibodies, carcinoembryonic antigen, streptococcus Group A antigen, viral antigens, antigens associated with autoimmune disease, allergens, tumor antigens, streptococcus Group B antigen, human immunodeficiency virus (HIV) I or HIV II antigen, antibodies, viruses, antigens specific to Rous sarcoma virus (RSV) antigen, enzyme, hormone, polysaccharide, protein, lipids, carbohydrate, drug, nucleic acid, Neisseria meningitides groups A, B, C, Y and W sub 135, Streptococcus pneumoniae, Escherichia coli K1, Haemophilus influenza type B, an antigen derived from microorganisms, a hapten, a drug of abuse, a therapeutic drug, an environmental agent, antigens specific to hepatitis, or especially Candida sp. or Salmonella sp. (claimed). The biosensors can be used to detect contamination in garments, such as diapers, and to detect contamination by microorganisms. They can also be used on contact lenses, eyeglasses, window panes, pharmaceutical vials, solvent containers, water bottles, or adhesive bandages, to detect contamination. ADVANTAGE - The method allows a modular production format so that Page 20

large rolls of patterned protein may be made for use with different analytes. The final product may be made by exposure to the necessary antibody. A diffraction image is produced which can be easily seen with the eye or, optionally, with a sensing device. The biosensors allow detection of extremely small quantities of analyte in a medium in a rapid assay lasting only a few minutes. In addition, no signaling or associate electronic components are required in the biosensing devices. Dwg.0/2

ANSWER 3 OF 7 WPIDS COPYRIGHT 2001 DERWENT INFORMATION LTD 1999-561863 [47] WPIDS ΑN DNC C1999-163794 Composition for inducing a type 2 T cell-independent immune response to TΙ an antigen, particularly for protection against bacteria. DC BO4 CO6 D16 MOND, J J; SNAPPER, C M IN (JACK-N) JACKSON FOUND ADVANCEMENT MILITARY MED PΑ CYC 21 A2 19990923 (199947) \* EN 41p PΙ WO 9947168 RW: AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE W: AU CA JP A 19991011 (200008) AU 9930054 WO 9947168 A2 WO 1999-US5647 19990315; AU 9930054 A AU 1999-30054 ADT 19990315 FDT AU 9930054 A Based on WO 9947168 19980316 PRAI US 1998-39247 9947168 A UPAB: 19991116 AB NOVELTY - Composition (A) comprises a type 2 T cell-independent antigen (Ag) conjugated to a lipid (or lipid-containing component) (B) that promotes an immune response to Ag. DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

(1) pharmaceutical composition or vaccine containing (A) dissolved

or

suspended in a carrier; and

(2) assay system for identifying compositions that stimulate isotype switching and/or a B cell memory response. ACTIVITY - Antibacterial.

MECHANISM OF ACTION - Induction of a specific immune response.

USE - (A) are used to promote a type 2 T cell-independent immune response to Ag (particularly a bacterial polysaccharide), especially in immunocompromised humans, e.g. patients deficient in T cell, neonates,

the

elderly, or subjects who are immunodefective because of exposure to viruses or other microorganisms, radiation, cytotoxic chemicals, corticosteroids or other immunosuppressants. (A) are useful in human or veterinary medicine.

ADVANTAGE - (A) elicit a primary response that is about 10 times greater than that produced when Ag and B are administered as separate components. They also induce a secondary response, i.e. a vigorous memory response and/or promotion of isotype switching. (A) are effective even in absence of functional cytokine-producing T cells. Mice were rendered T cell defective by administration of an anti-CD4 antibody, then one day later they were injected with 5 micro g of (i) Streptococcus pneumoniae type 14 polysaccharide/lipoprotein D complex or (ii) the polysaccharide only. After 14 days, the immunoglobulin G1 titer (ng/ml)

against the polysaccharide was 5834 in (i) but less than 10 in (ii).

Dwg.0/0

ANSWER 4 OF 7 WPIDS COPYRIGHT 2001 DERWENT INFORMATION LTD WPIDS 1999-443884 [37] ANDNC C1999-130702 DNN N1999-331070 Biosensors used to detect analyte, e.g. chemical or biological TΙ contamination in garments, e.g. diapers. DC A18 A23 A26 A96 B04 D16 D22 E13 J04 P75 S03 EVERHART, D S; JONES, M L; KAYLOR, R M ΙN (KIMB) KIMBERLY-CLARK WORLDWIDE INC PA CYC 84 A1 19990624 (199937)\* EN WO 9931486 ·39p PΙ RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL OA PT SD SE SZ UG ZW W: AL AM AT AU AZ BA BB BG BR BY CA CH CN CU CZ DE DK EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT UA UG UZ VN YU ZW AU 9919205 Α 19990705 (199948) A 20000509 (200030) A1 20001004 (200050) US 6060256 EN EP 1040338 R: BE DE ES FR GB IT NL SE A 20010307 (200140) CN 1286753 ADT WO 9931486 A1 WO 1998-US26759 19981216; AU 9919205 A AU 1999-19205 19981216; US 6060256 A US 1997-991644 19971216; EP 1040338 A1 EP 1998-963991 19981216, WO 1998-US26759 19981216; CN 1286753 A CN 1998-812255 19981216 FDT AU 9919205 A Based on WO 9931486; EP 1040338 Al Based on WO 9931486 PRAI US 1997-991644 19971216 9931486 A UPAB: 19990914 NOVELTY - Biosensors comprising: (a) polymer film coated with metal; and (b) patterned receptor layer printed onto (a) on which is a receptive material that specifically binds analyte. DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for: (1) Methods of detecting analyte using the sensor; and (2) A method of making the biosensor. USE - Used to detect analyte. Used particularly in the field of microcontact printing binders on metal films to produce optical diffraction biosensors. Used as single tests for detecting analyte or as multiple test devices. Used for detection of chemical or biological contamination in garments such as diapers, detection of contamination by microorganisms in pre-packed foods such as fruit juices and other beverages, and in health diagnostic applications such as diagnostic kits for detection of antigens, microorganisms, and blood constituents. May be used on contact lenses, eyeglasses, windowpanes, pharmaceutical vials, solvent containers, water bottles and plasters to detect contamination. Used in immunoassays for either antigen or antibody detection, for use in direct, indirect or competitive detection systems, for determination of enzymatic activity, for detection of small organic molecules (drugs of abuse, therapeutic drugs, environmental agents) and nucleic acids. with patterned antibody to Candida albicans were prepared by

pretreating gold/polyester (10 nm thick) by immersion in 5 mg/ml

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phosphate-buffered saline solution (pH 7.2) of beta-casein for 10
minutes.
    The sample was rinsed thoroughly with distilled water and dried under a
    strong nitrogen stream. Contact printing was done using a
    polydimethylsiloxane stamp with an x,y array of 10 mu m diameter circles.
    The stamp was coated with a thiolated antibody to Candida
    albicans by immersing in a 0.5 mg/ml aqueous solution of antibody
    derivative. After 10 minutes, the stamp was removed and thoroughly dried
    using a strong stream of nitrogen. Contact printing was done on the
     casein-treated sample, with exposure times of 1 second to 2 minutes being
     adequate. After printing, the sample was again rinsed with distilled
    and dried. The sensor sample was exposed to germ tube-bearing cells of C.
    albicans by inoculating tape-stripped adult forearm skin with a
    concentration of 106 yeast cells/ml and placing the sensor on top of the
    yeast-containing tape. Transfer of the yeast cells to the sensor was
    accomplished after only a few seconds of contact. Patterned adhesion of
    the yeast cells to the sensor was confirmed by microscopic analysis and
     resulted in a diffraction image upon irradiation with a laser.
         ADVANTAGE - Are inexpensive and sensitive devices. Produced by easy,
     efficient and simple method of contact printing a patterned receptor on
an
    optically transparent, flexible substrate, that is amenable to continuous
    processing and does not use self-assembling monolayers. Are simpler than
    prior art, are not restricted to limitations of self-assembling
monolayers
    and are easier to manufacture. Are low-cost and disposable and can be
    mass-produced.
          DESCRIPTION OF DRAWING(S) - Schematic representation of metal-plated
    MYLAR (RTM) film with nutrient backing.
         MYLAR film (RTM: polyethylene-terephthalate) 15
    metal film 20
          receptors specific for microorganism 25
          nutrient backing 30
     Dwg.1/8
                                           DERWENT INFORMATION LTD
    ANSWER 5 OF 7 WPIDS COPYRIGHT 2001
L19
     1997-340536 [31]
                       WPIDS
ΑN
                       1991-117625 [16]; 1992-300174 [36]; 1992-300183
CR
     1991-117618 [16];
[36];
     1992-349359 [42]; 1994-065810 [08]; 1996-010090 [01]; 1996-076885
[08];
     1996-361950 [36]; 1997-288174 [26]; 1999-152762 [13]
                       DNC C1997-109335
DNN
    N1997-282600
     Optimisation of visual signal from optical assay device - for
ΤI
     detection of analyte, e.g. rheumatoid factor, viral antigens,
     carbohydrate, drug or nucleic acid.
DC
     B04 D16 J04 S02 S03
     BOGART, G R; ETTER, J B
ΙN
     (BIOS-N) BIOSTAR INC
PΑ
CYC
                   A 19970617 (199731)*
PΙ
     US 5639671
                                              69p
    US 5639671 A CIP of US 1989-408291 19890918, CIP of US 1992-873097
ADT
     19920424, CIP of US 1992-923048 19920731, Cont of US 1993-76319 19930610,
     US 1995-412600 19950328
                      19930610; US 1989-408291
                                                 19890918; US 1992-873097
PRAI US 1993-76319
                              19920731; US 1995-412600
                                                           19950328
     19920424; US 1992-923048
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5639671 A UPAB: 19990331
AB
    Optimizing a visual signal from an optical assay device for the
     detection of an analyte comprises: (a) providing a substrate having an
     anti-reflective film on it, with a series of thicknesses (preferably
     optimal) varied incrementally along the length of the substrate,
providing
     an attachment layer of a chosen thickness on the anti-reflective film and
     providing a receptive layer of a chosen thickness for the analyte on the
     attachment layer; (b) contacting the analyte with the receptive layer
such
     that a mass change on the receptive layer results, and (c) determining at
     least 1 thickness of the series of thicknesses of the anti-reflective
     that, in combination with the other layers of the device, produces a
     visual signal comprising a maximised visual contrast in interference
     colour upon the change in mass relative to a background interference
     colour, over a range of concentrations of the analyte.
          USE - The method can be used to detect analytes such as rheumatoid
     factor, immunoglobulin E antibodies specific for Birch pollen,
     carcinoembryonic antigen, Streptococcus Group A antigen
     , viral antigens, antigens associated with autoimmune
     disease, allergens, a tumour or an infectious microorganism,
Streptococcus
     Group B antigen, HIV I or HIV II antigen, host
     response (antibodies) to the virus, antigens specific
     to RSV or host response (antibodies) to the virus, an
     antibody, antigen, enzyme, hormone,
    polysaccharide, protein, lipid, carbohydrate, drug or nucleic
     acid, analyte derived from causative organisms for meningitis, Neisseria
     meningitidis groups A, B, C, Y and W135, S. pneumoniae, E. coli K1,
    Haemophilus influenzae type B, antigen derived from
    microorganisms, a hapten, a drug of abuse (including drugs which are
     unlawful to use without a permit or license), a therapeutic drug, an
     environmental agent and antigens specific to hepatitis.
     Dwg.0/18
    ANSWER 6 OF 7 WPIDS COPYRIGHT 2001
                                           DERWENT INFORMATION LTD
L19
     1994-200269 [24]
                        WPIDS
AN
DNC
    C1994-091569
     Nucleic acid encoding D15 outer membrane protein - esp. of
     Haemophilus influenzae, and related proteins, vectors, antisera
     etc. useful in vaccines, for diagnosis and for passive immunisation..
     B04 D16
IN
     CHONG, P; KLEIN, M; LOOSMORE, S; SIA, D Y C; THOMAS, W; YANG, Y;
LOOSMORE,
     S M; SIA, D; YANG, Y P
     (CONN-N) CONNAUGHT LAB LTD
PA
CYC
     28
                   A1 19940609 (199424)*
ΡI
     WO 9412641
                                             161p
        RW: AT BE CH DE DK ES FR GB GR IE IT LU MC NL PT SE
         W: AU BR CA FI JP KR NO NZ RU UA US
                   A 19940622 (199436)
     AU 9455565
                   A1 19950830 (199539)
                                        ΕN
     EP 668916
         R: AT BE CH DE DK ES FR GB GR IE IT LI LU MC NL PT SE
                  W 19960319 (199644)
                                             156p
     JP 08502417
                  B 19971113 (199803)
     AU 683435
     BR 9307510 A 19990601 (199927)
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JP 2907552
                                             181p
                   B2 19990621 (199930)
                   A 20000111 (200010)
    US 6013514
                   A 20000704 (200036)
    US 6083743
    RU 2141528
                   C1 19991120 (200041)
                   B1 19990816 (200104)
    KR 216390
    WO 9412641 A1 WO 1993-CA501 19931123; AU 9455565 A AU 1994-55565
19931123;
    EP 668916 A1 WO 1993-CA501 19931123, EP 1994-900671 19931123; JP 08502417
    W WO 1993-CA501 19931123, JP 1994-512608 19931123; AU 683435 B AU
    1994-55565 19931123; BR 9307510 A BR 1993-7510 19931123, WO 1993-CA501
     19931123; JP 2907552 B2 WO 1993-CA501 19931123, JP 1994-512608 19931123;
    US 6013514 A WO 1993-CA501 19931123, US 1995-433522 19950912; US 6083743
    Cont of WO 1993-CA501 19931123, Cont of US 1995-433522 19950912, US
    1998-135166 19980818; RU 2141528 C1 WO 1993-CA501 19931123, RU
1995-117238
     19931123; KR 216390 B1 WO 1993-CA501 19931123, KR 1995-702081 19950523
FDT AU 9455565 A Based on WO 9412641; EP 668916 Al Based on WO 9412641; JP
    08502417 W Based on WO 9412641; AU 683435 B Previous Publ. AU 9455565,
    Based on WO 9412641; BR 9307510 A Based on WO 9412641; JP 2907552 B2
    Previous Publ. JP 08502417, Based on WO 9412641; US 6013514 A Based on WO
     9412641; RU 2141528 Cl Based on WO 9412641
PRAI GB 1992-24584
                      19921123
          9412641 A UPAB: 19940803
    WO
    New nucleic acid (I) contains at least a portion coding for a D15 outer
    membrane protein (omp) and has a sequence which is (a) any of 5 (all
about
    3000bp) reproduced in the specification, or complementary sequences or
(b)
    hybridsable under stringent conditions with such sequences. Also new are
    (1) recombinant plasmids contg. a segment of (I) at least 18 bp long (and
    opt. expression control elements, (12) proteins (II) encoded by these
    plasmids; (3) purified D15 omp (III); (4) synthetic polypeptides with
     sequences corresp. to (II) or (III), or their variants and mutants which
    retain immunogenicity; (5) antisera or antibodies specific for
     (II), (III) or immunologous contg. them; (6) chimeric molecules
consisting
     of (II) or (III) bonded to another polypeptides, protein or
    polysaccharides.
          USE - (I), (II) and the synthetic polypeptides are useful in
vaccines
    to protect against Haemophilus. D15 can also be used as a
    carrier for polysaccharide antigens to form conjugate
    vaccines against other bacteria; to induce immunity to abnormal
    polysaccharides or tumour cells and to generate anti-tumour
     antibodies, for coupling to toxins etc. (I), (II) synthetic
    peptides and antisera can also be used diagnostically (in hybridisation
or
     immunoassay procedures) and antibodies can be used for
     passive immunisation.
     Dwg.0/11
    ANSWER 7 OF 7 WPIDS COPYRIGHT 2001
                                           DERWENT INFORMATION LTD
     1986-292987 [45]
                        WPIDS
AN
     1991-209899 [29]
CR
                        DNC C1986-126914
DNN
    N1986-218851
     Solid phase system for use in ligand-receptor assays - e.g. for
TI
                                                                       Page 25
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detection of human choriogonadotropin antigen in urine.
DC
    A89 B04 D16 J04 S03
IN
    RUBENSTEIN, A S
     (HYBR-N) HYBRITECH INC
PA
CÝC
    18
                                              30p
                   A 19861105 (198645) * EN
PΙ
        R: AT BE CH DE FR GB IT LI LU NL SE
                   A 19861016 (198648)
    AU 8655657
    NO 8601317
                   Α
                     19861027 (198650)
                   A 19861005 (198703)
    FI 8601460
                   A 19861222 (198705)
    JP 61292059
                   A 19861005 (198706)
    DK 8601538
                   A 19890501 (198924)
    ES 8900176
    CA 1272127
                   A 19900731 (199036)
                   B1 19930127 (199304)
                                              14p
    EP 200381
                                         EN
         R: AT BE CH DE FR GB IT LI LU NL SE
                   G 19930311 (199311)
    DE 3687589
                   B 19940630 (199428)
    FI 92257
                   B2 19951206 (199602)
                                              13p
     JP 07113636
                   B 19970811 (199739)
     DK 171928
    EP 200381 A EP 1986-302521 19860404; JP 61292059 A JP 1986-78064
19860404;
     ES 8900176 A ES 1986-553724 19860404; EP 200381 B1 EP 1986-302521
     19860404; DE 3687589 G DE 1986-3687589 19860404, EP 1986-302521 19860404;
     FI 92257 B FI 1986-1460 19860404; JP 07113636 B2 JP 1986-78064 19860404;
     DK 171928 B DK 1986-1538 19860404
    DE 3687589 G Based on EP 200381; FI 92257 B Previous Publ. FI 8601460; JP
FDT
     07113636 B2 Based on JP 61292059; DK 171928 B Previous Publ. DK 8601538
PRAI US 1985-720036
                      19850404
           200381 A UPAB: 19940907
    EΡ
     Solid phase system for use in a ligand-receptor assay for the
     detection of a selected analyte in a fluid sample comprises a porous
    matrix in which microspheres are entrapped, the microspheres being bound
    with a receptor capable of capturing a target ligand.
          The system is for use in a multiple ligand-receptor assay
     for simultaneous detection of at least two selected analytes, distinct
     gps. of microspheres that are entrapped within discrete zones in the
     porous matrix, each gp. of microspheres being bound with a receptor
     capable of capturing a different target ligand.
          The following specific cases are claimed: one gp. of microspheres is
     bound with a receptor capable of capturing a target ligand; and one gp.
οf
     microspheres is bound with either (1) the target ligand or other receptor
     substance as a positive control; or (2) a substance incapable of
capturing
     the target ligand without a bound component, as a negative control for
     detection the analyte. Appts. is also provided.
          USE/ADVANTAGE - The system is useful for assay of
     antigens such as prostatic acid phosphatase, prostate-specific
     antigen alpha foetoprotein, carcinoembryonic antigen,
     lentenising hormone and creatine kinase isoenzyme. It is also useful for
     the assay of viruses, bacteria, parasites or fungi or associated
     antigens or antibodies, e.g. rubella, rota, hepatitis,
     herpes and influenza viruses, HTLV, group A and group B streptococcus,
     Neisseria gonorrhea, Trichomonas vaginalis, Candida albicans, Chamydia
     trachomatis and Haemophilus influenza. The system may also be
                                                                        Page 26
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employed in nucleic acid probe technology. 0/6 Dwg. 0/6

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#### => fil medline

FILE 'MEDLINE' ENTERED AT 14:58:00 ON 06 AUG 2001

FILE LAST UPDATED: 30 JUL 2001 (20010730/UP). FILE COVERS 1958 TO DATE.

On April 22, 2001, MEDLINE was reloaded. See HELP RLOAD for details.

MEDLINE now contains new records from the former NLM HEALTH STAR database. These records have an Entry Date and Update Date of 20010223.

MEDLINE thesauri in the /CN, /CT, and /MN fields incorporate the MeSH 2001 vocabulary. Enter HELP THESAURUS for details.

The OLDMEDLINE file segment now contains data from 1958 through 1965. Enter HELP CONTENT for details.

Left, right, and simultaneous left and right truncation are available in the Basic Index. See HELP SFIELDS for details.

THIS FILE CONTAINS CAS REGISTRY NUMBERS FOR EASY AND ACCURATE SUBSTANCE IDENTIFICATION.

#### => d his

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(FILE 'MEDLINE' ENTERED AT 14:51:27 ON 06 AUG 2001)
                DEL HIS Y
            226 S IMMUNOCHROMATOGRA?
L1
                E HAEMOPHILUS/CT
           2001 S HAEMOPHILUS/CT
1.2
              0 S L1 AND L2
L3
                E IMMUNOCHEMISTRY/CT
                E E3+AL
                E E3+ALL
         258309 S IMMUNOCHEMISTRY+NT/CT
L4
                E CHROMATOGRAPHY/CT
                E E3+ALLK
                E E3+ALL
              0 S CHROMATOGRAPY/CT
L_5
          26123 S CHROMATOGRAPHY/CT
L6
              7 S L6 AND L2
L7
L8
              0 S L7 AND L4
L9
          10860 S ASSAY/CT OR IMMUNOASSAY/CT
L10
          10860 S ASSAYS/CT OR IMMUNOASSAY/CT
              0 S L10 AND L7
L11
              2 S L7 AND (ASSAY? OR IMMUNOASSAY?)
L12
          99901 S STRIP# OR APP## OR APPARATUS
L13
             12 S L13 AND L2
L14
              0 S L14 AND (L6 OR L10)
L15
                E GOLD/CT
           6189 S GOLD/CT
L16
              1 S L16 AND L2
L17
             15 S L12 OR L14 OR L17
L18
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FILE 'MEDLINE' ENTERED AT 14:58:00 ON 06 AUG 2001

#### => d .med 118 1-15

L18 ANSWER 1 OF 15 MEDLINE 2000087378 MEDLINE AN DN PubMed ID: 10618273 Immunoglobulins in nasal secretions of healthy humans: structural ΤI integrity of secretory immunoglobulin A1 (IgA1) and occurrence of neutralizing antibodies to IgAl proteases of nasal bacteria. Kirkeby L; Rasmussen T T; Reinholdt J; Kilian M ΑU Department of Medical Microbiology and Immunology, University of Aarhus, CS DK-8000 Aarhus C, Denmark. CLINICAL AND DIAGNOSTIC LABORATORY IMMUNOLOGY, (2000 Jan) 7 (1) 31-9. SO Journal code: CB7; 9421292. ISSN: 1071-412X. CY United States Journal; Article; (JOURNAL ARTICLE) DT LA English Priority Journals FS EΜ 200003 Entered STN: 20000314 ED Last Updated on STN: 20000314 Entered Medline: 20000302 Certain bacteria, including overt pathogens as well as commensals, AΒ produce immunoglobulin A1 (IgA1) proteases. By cleaving IgA1, including secretory IgA1, in the hinge region, these enzymes may interfere with the barrier functions of mucosal IgA antibodies, as indicated by experiments in Previous studies have suggested that cleavage of IgAl in nasal secretions may be associated with the development and perpetuation of atopic disease. To clarify the potential effect of IgAl protease-producing bacteria in the nasal cavity, we have analyzed immunoglobulin isotypes in nasal secretions of 11 healthy humans, with a focus on IgA, and at the same time have characterized and quantified IgA1 protease-producing bacteria in the nasal flora of the subjects. Samples in the form of nasal wash were collected bу using a washing liquid that contained lithium as an internal reference. Dilution factors and, subsequently, concentrations in undiluted secretions could thereby be calculated. IgA, mainly in the secretory form, was found by enzyme-linked immunosorbent assay to be the dominant isotype in all subjects, and the vast majority of IgA (median, 91%) was of the Al subclass, corroborating results of previous analyses at the level of immunoglobulin-producing cells. Levels of serum-type immunoglobulins were low, except for four subjects in whom levels of IgG corresponded to 20 to

66% of total IgA. Cumulative levels of IgA, IgG, and IgM in undiluted secretions ranged from 260 to 2,494 (median, 777) &mgr;g ml(-1). IgAl protease-producing bacteria (Haemophilus influenzae, Streptococcus

undiluted secretion, corresponding to 0.2 to 99.6% of the flora.

pneumoniae, or Streptococcus mitis biovar 1) were isolated from the nasal cavities of seven subjects at  $2.1 \times 10(3)$  to  $7.2 \times 10(6)$  CFU per ml of

Nevertheless, alpha-chain fragments characteristic of IgAl protease
Page 29

activity were not detected in secretions from any subject by immunoblotting. Neutralizing antibodies to IgAl proteases of autologous isolates were detected in secretions from five of the seven subjects but not in those from two subjects harboring IgA1 protease-producing S. mitis biovar 1. alpha-chain fragments different from Fc(alpha) and Fd(alpha) were detected in some samples, possibly reflecting nonspecific proteolytic activity of microbial or host origin. These results add to previous evidence for a role of secretory immunity in the defense of the nasal mucosa but do not help identify conditions under which bacterial IgA1 proteases may interfere with this defense. CTCheck Tags: Human; Support, Non-U.S. Gov't \*Antibodies: IM, immunology Bacteria: EN, enzymology \*Bacteria: IM, immunology Child Chromatography Haemophilus: ME, metabolism \*IgA: CH, chemistry \*IqA, Secretory: CH, chemistry Immunoblotting \*Immunoglobulins: AN, analysis Middle Age \*Nasal Mucosa: IM, immunology Neutralization Tests Protease Inhibitors: PD, pharmacology \*Serine Endopeptidases: IM, immunology Streptococcus: ME, metabolism ANSWER 2 OF 15 MEDLINE - T.18 MEDLINE AN 1998395109 DN 98395109 PubMed ID: 9727006 Purification and characterization of wild-type and mutant "classical" TInitroreductases of Salmonella typhimurium. L33R mutation greatly diminishes binding of FMN to the nitroreductase of S. typhimurium. Watanabe M; Nishino T; Takio K; Sofuni T; Nohmi T ΑU Division of Genetics and Mutagenesis, National Institute of Health CS Sciences, 1-18-1 Kamiyoga, Setagaya-ku, Tokyo 158-8501, Japan. JOURNAL OF BIOLOGICAL CHEMISTRY, (1998 Sep 11) 273 (37) 23922-8. SO Journal code: HIV; 2985121R. ISSN: 0021-9258. CY United States  $\mathsf{D}\mathbf{T}$ Journal; Article; (JOURNAL ARTICLE) LA English FS Priority Journals EM199810 ED Entered STN: 19981021 Last Updated on STN: 19981021 Entered Medline: 19981013 "Classical" nitroreductase of Salmonella typhimurium is a flavoprotein AΒ that catalyzes the reduction of nitroaromatics to metabolites that are toxic, mutagenic, or carcinogenic. This enzyme represents a new class of flavin-dependent enzymes, which includes nitroreductases of Enterobacter cloacae and Escherichia coli, flavin oxidoreductase of Vibrio fischeri, and NADH oxidase of Thermus thermophilus. To investigate the structure-function relation of this class of enzymes, the gene encoding a mutant nitroreductase was cloned from S. typhimurium strain TA1538NR, and

the enzymatic properties were compared with those of the wild-type. DNA sequence analysis revealed a T to G mutation in the mutant nitroreductase gene, predicting a replacement of leucine 33 with arginine. In contrast to the wild-type enzyme, the purified protein with a mutation of leucine 33 to arginine has no detectable nitroreductase activities in the standard assay conditions and easily lost FMN by dialysis or ultrafiltration. In the presence of an excess amount of FMN, however, the mutant protein exhibited a weak but measurable enzyme activity, and the substrate specificity was similar to that of the wild-type enzyme. Possible mechanisms by which the mutation greatly diminishes binding of FMN to the nitroreductase are discussed. Check Tags: Comparative Study; Support, Non-U.S. Gov't CT Amino Acid Sequence Base Sequence Binding Sites Chromatography Chromatography, Gel Chromatography, Ion Exchange Cloning, Molecular Durapatite Enterobacter cloacae: EN, enzymology Escherichia coli: EN, enzymology \*FMN: ME, metabolism Haemophilus: EN, enzymology Kinetics Molecular Sequence Data Molecular Weight Mycoplasma: EN, enzymology \*Nitroreductases: GE, genetics Nitroreductases: IP, isolation & purification \*Nitroreductases: ME, metabolism \*Point Mutation \*Salmonella typhimurium: EN, enzymology Salmonella typhimurium: GE, genetics Sequence Alignment Sequence Homology, Amino Acid Thermus thermophilus: EN, enzymology Vibrio: EN, enzymology ANSWER 3 OF 15 MEDLINE L18 MEDLINE AN 95298926 PubMed ID: 7779956 DN 95298926 Evaluation of a commercial system for the identification of nonfermenting bacteria of veterinary importance. Salmon S A; Watts J L; Walker R D; Yancey R J Jr ΑU Animal Health Discovery Research, Upjohn Company, Kalamazoo, MI 49001, CS JOURNAL OF VETERINARY DIAGNOSTIC INVESTIGATION, (1995 Jan) 7 (1) 161-4. SO Journal code: A2D; 9011490. ISSN: 1040-6387. CY United States Journal; Article; (JOURNAL ARTICLE) DT

LA

FS

EΜ

English

199507

Priority Journals

Entered STN: 19950726

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Last Updated on STN: 19950726
     Entered Medline: 19950718
    Check Tags: Animal
     *Bacteriological Techniques: VE, veterinary
     *Gram-Negative Bacteria: CL, classification
     *Gram-Negative Bacteria: IP, isolation & purification
     *Gram-Negative Facultatively Anaerobic Rods: CL, classification
     *Gram-Negative Facultatively Anaerobic Rods: IP, isolation & purification
     Haemophilus
     Reagent Strips
    ANSWER 4 OF 15
                        MEDLINE
     92112340
                  MEDLINE
AN
                PubMed ID: 1730505
DN
     92112340
    Clustering of an outer membrane adhesin of Haemophilus parainfluenzae.
TΙ
     Liljemark W F; Bloomquist C G; Lai C H
ΑU
     Department of Diagnostic and Surgical Sciences, School of Dentistry,
CS
     University of Minnesota, Minneapolis 55455-0329.
NC
     DE07014 (NIDCR)
     R37-DE04614 (NIDCR)
     INFECTION AND IMMUNITY, (1992 Feb) 60 (2) 687-9.
SO
     Journal code: GO7; 0246127. ISSN: 0019-9567.
CY
     United States
     Journal; Article; (JOURNAL ARTICLE)
DT
LA
     English
     Priority Journals
FS
     199202
EM
     Entered STN: 19920308
ED
     Last Updated on STN: 20000303
     Entered Medline: 19920218
     Haemophilus parainfluenzae synthesizes an outer membrane protein adhesin
AB
     which mediates binding to oral streptococci, salivary pellicle, and
     neuraminidase-treated erythrocytes. An indirect gold labeling technique
     and immunoelectron microscopy verified the location of this outer
membrane
     protein. Further, a clustering of gold particles was observed in
irregular
     patches at the cell surface.
     Check Tags: Support, U.S. Gov't, P.H.S.
CT
     *Bacterial Adhesion
     *Bacterial Outer Membrane Proteins: AN, analysis
      Gold
     *Haemophilus: CH, chemistry
L18
     ANSWER 5 OF 15
                        MEDLINE
AN
     91263343
                  MEDLINE
                PubMed ID: 2048282
DN
     91263343
     Improved protection of swine from pleuropneumonia by vaccination with
TI
     proteinase K-treated outer membrane of Actinobacillus (Haemophilus)
     pleuropneumoniae.
     Chiang Y W; Young T F; Rapp-Gabrielson V J; Ross R F
ΑU
     Veterinary Medical Research Institute, College of Veterinary Medicine,
CS
     Iowa State University, Ames 50011.
     VETERINARY MICROBIOLOGY, (1991 Mar) 27 (1) 49-62.
SO
     Journal code: XBW; 7705469. ISSN: 0378-1135.
     Netherlands
CY
     Journal; Article; (JOURNAL ARTICLE)
DT
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LA

English

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Priority Journals
FS
EM
     199107
     Entered STN: 19910802
F.D
     Last Updated on STN: 20000303
     Entered Medline: 19910718
     The immunogenic and protective potentials of an outer membrane-enriched
AΒ
     fraction (OM) from a serotype 5 strain of Actinobacillus (Haemophilus)
    pleuropneumoniae (APP) and the same OM degraded with proteinase
     K or periodate were evaluated in swine. Groups of pigs were vaccinated
    with two doses of OM, proteinase K-treated OM (P-OM), periodate-treated
MO
     (PI-OM), or placebo vaccine and challenged intranasally with the
    homologous strain of APP. Results from triplicate experiments
     indicated that proteinase K treatment of OM resulted in an improved
     efficacy. This improved efficacy of P-OM vaccine over untreated OM
vaccine
    was evidenced not only by less severe lung lesions in P-OM vaccinated
pigs
    but also by significant reduction (P less than 0.05) in the number of
P-OM
    vaccinated pigs which developed lung lesions upon challenge with
    APP. Assessment of sera from vaccinated animals by immunoblotting,
     complement fixation test, or ELISA indicated that the immunogenicity of
     some but not all protein or carbohydrate components were reduced (or
     eliminated) by proteinase K and periodate treatments respectively.
    Check Tags: Animal; Female; Male; Support, Non-U.S. Gov't
CT
     *Actinobacillus: IM, immunology
     Actinobacillus Infections: PC, prevention & control
     *Actinobacillus Infections: VE, veterinary
      Antibodies, Bacterial: BI, biosynthesis
      Antigens, Bacterial: IM, immunology
      Bacterial Outer Membrane Proteins: IM, immunology
      Complement Fixation Tests
      Electrophoresis, Polyacrylamide Gel
      Endopeptidase K
      Enzyme-Linked Immunosorbent Assay
      Haemophilus: IM, immunology
      Haemophilus Infections: PC, prevention & control
      Haemophilus Infections: VE, veterinary
      Pleuropneumonia: PC, prevention & control
     *Pleuropneumonia: VE, veterinary
      Random Allocation
      Serine Endopeptidases: ME, metabolism
     *Swine Diseases: PC, prevention & control
     *Vaccination: VE, veterinary
    ANSWER 6 OF 15
                        MEDLINE
L18
                 MEDLINE
     88047453
AN
                PubMed ID: 3674414
     88047453
DN
     Preparation of cell envelopes of large numbers of individual bacterial
TI
     strains with the use of an automatic cell disruptor.
     Van Alphen L; Romijn C; Brandt H; Geelen L; Zanen H C
ΑU
     Department of Medical Microbiology, Academic Medical Centre, Amsterdam,
CS
     The Netherlands.
     ANALYTICAL BIOCHEMISTRY, (1987 Oct) 166 (1) 36-40.
SO
                                                                        Page 33
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Journal code: 4NK; 0370535. ISSN: 0003-2697. CY United States Journal; Article; (JOURNAL ARTICLE) DT LA English FS Priority Journals EM 198711 Entered STN: 19900305 ED Last Updated on STN: 19900305 Entered Medline: 19871124 Analysis of the cell envelopes of large numbers of bacterial strains is AΒ used for the epidemiological and taxonomic investigation of clinical, veterinarian, and ecological isolates. Isolation of cell envelopes requires lysis of the bacteria. We developed an apparatus to disrupt bacterial cells of 200 different isolates in suspension by ultrasonication automatically. It is composed of modified standard laboratory equipment (fraction collector, cooling unit, pump), a standard ultrasonifier, and a newly designed control unit, which includes a sampler. This apparatus was applied to the analysis of cell envelope proteins of 96 Haemophilus influenzae strains on sodium dodecyl sulfate-polyacrylamide gel electrophoresis within 3 days after the first culture. \*Bacterial Outer Membrane Proteins: AN, analysis CTChemistry, Analytical: IS, instrumentation Electrophoresis, Polyacrylamide Gel Haemophilus: AN, analysis L18 ANSWER 7 OF 15 MEDLINE 87132255 MEDLINE AN 87132255 PubMed ID: 3493113 DN Rapid determination of X/V growth requirements of Haemophilus species in ΤI broth. Inzana T J; Clarridge J; Williams R P ΑU AI07145 (NIAID) NC DIAGNOSTIC MICROBIOLOGY AND INFECTIOUS DISEASE, (1987 Feb) 6 (2) 93-100. SO Journal code: DMI; 8305899. ISSN: 0732-8893. CY United States Journal; Article; (JOURNAL ARTICLE) DTLA English FS Priority Journals EΜ 198704 Entered STN: 19900303 Last Updated on STN: 19970203 Entered Medline: 19870423 A broth system was developed for rapid identification of the requirement for X factor (hemin), or V factor (NAD), or both for growth of Haemophilus species. This system was compared to growth around paper discs/ strips impregnated with factors X and/or V. The broth system consisted of three tubes, each containing brain-heart infusion broth supplemented with V factor, X factor, or both. Each tube was inoculated with a saline suspension of an Haemophilus isolate, and the broths were shaken for aeration at 37 degrees C. Under these conditions turbidity or clumping was usually evident after 4-5 hr only in the broth(s) containing the required supplement(s). A few strains requiring only V factor required

overnight incubation. One hundred fifty-six Haemophilus isolates were

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tested for growth around supplemented discs/strips or in
    supplemented broths: 129 were H. influenzae/aegypticus, 25 were of
    species that required only V factor, and 2 were H. aphrophilus. Ten of 89
    H. influenzae isolates from the respiratory tract were misidentified by
    satellitism. All isolates were correctly identified by growth in
    supplemented broths. The cost of the broth assay was about 60 cents/test,
    whereas the satellite assay cost about 120 cents/test. Serotyping and
    antibiotic sensitivity testing could be performed directly from the broth
    culture. Determination of X and/or V requirement by Haemophilus species
    with supplemented broths was sensitive, rapid, and inexpensive.
    Check Tags: Animal; Comparative Study; Human; Support, U.S. Gov't, P.H.S.
     Blood: MI, microbiology
     Cerebrospinal Fluid: MI, microbiology
     Costs and Cost Analysis
     Culture Media
     Ear: MI, microbiology
     Eye: MI, microbiology
     *Haemophilus: GD, growth & development
     Haemophilus: IP, isolation & purification
     *Haemophilus influenzae: GD, growth & development
      Haemophilus influenzae: IP, isolation & purification
     *Heme: AA, analogs & derivatives
     *Hemin: PD, pharmacology
      Kinetics
     *NAD: PD, pharmacology
     Respiratory System: MI, microbiology
L18 ANSWER 8 OF 15
                        MEDLINE
                MEDLINE
    86168818
AΝ
               PubMed ID: 3514664
DN
    86168818
    Comparison of a new commercially prepared porphyrin test and the
ТΤ
     conventional satellite test for the identification of Haemophilus species
     that require the X factor.
ΑU
    Gadberry J L; Amos M A
     JOURNAL OF CLINICAL MICROBIOLOGY, (1986 Mar) 23 (3) 637-9.
SO
     Journal code: HSH; 7505564. ISSN: 0095-1137.
CY
    United States
DT
     Journal; Article; (JOURNAL ARTICLE)
LA
    English
     Priority Journals
FS
EM
    198605
ED
    Entered STN: 19900321
    Last Updated on STN: 19900321
    Entered Medline: 19860514
    A test with a commercially developed porphyrin test agar was examined for
AB
    the identification of Haemophilus spp. The porphyrin test agar method was
     compared with the conventional paper strip satellite method in
     tests with 187 isolates and was found to be easier to perform and
     interpret, giving a sensitivity of 98.7% and specificity of 94.7%.
CT
    Check Tags: Comparative Study
      Agar
     *Bacteriological Techniques
      Culture Media
     *Haemophilus: CL, classification
      Haemophilus: GD, growth & development
     Haemophilus: ME, metabolism
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\*Heme: AA, analogs & derivatives

\*Hemin: PD, pharmacology

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*Porphyrins: BI, biosynthesis
    ANSWER 9 OF 15
                        MEDLINE
L18
                  MEDLINE
    85073854
AN
    85073854
                PubMed ID: 6509390
DN
     Porcine haemophili and actinobacilli: characterization by means of API
ΤI
     test strips and possible taxonomic implications.
     O'Reilly T; Rosendal S; Niven D F
ΑU
    CANADIAN JOURNAL OF MICROBIOLOGY, (1984 Oct) 30 (10) 1229-38.
SO
     Journal code: CJ3; 0372707. ISSN: 0008-4166.
CY
    Canada
    Journal; Article; (JOURNAL ARTICLE)
DT
LA
    English
    Priority Journals
FS
    198502
ΕM
     Entered STN: 19900320
ED
    Last Updated on STN: 19900320
     Entered Medline: 19850221
    Thirty Haemophilus strains and six Actinobacillus strains, all of porcine
AΒ
     origin, were examined for their biochemical reactivity on API 20E and API
     ZYM test strips using dense cell suspensions (supplemented with
     NAD as appropriate) as strip inocula. When combined with a test
     for V-factor dependency, the use of both strips allowed adequate
     differentiation of closely related organisms. Numerical taxonomic
     of the data demonstrated that the majority of the haemophili and
     actinobacilli studied could be placed in one of four major clusters;
     clusters contained, respectively, the H. pleuropneumoniae--A.
     pleuropneumoniae strains, the H. parasuis strains, strains belonging to
     Haemophilus taxon "minor group," and strains belonging to an unusual
group
     of mannitol-positive, urease-negative haemophili. A representative of
     Haemophilus species taxon C and an unusual Actinobacillus isolate
     to be comparatively unrelated to organisms in the four major clusters.
     Although it may, on occasion, be difficult to place an unusual isolate in
     any one particular group, owing to the uncertain taxonomy of some of
these
     organisms, it is concluded that API test strips can serve as
     useful tools for the characterization and differentiation of porcine
     haemophili and actinobacilli.
     Check Tags: Animal; Support, Non-U.S. Gov't
CT
     *Actinobacillus: CL, classification
     Actinobacillus: EN, enzymology
     *Haemophilus: CL, classification
     Haemophilus: EN, enzymology
     *Indicators and Reagents
     *Reagent Strips
     *Swine: MI, microbiology
L18 ANSWER 10 OF 15
                         MEDLINE
     84207971
                  MEDLINE
ΑN
                PubMed ID: 6373253
DN
     A modified method for differentiation of Haemophilus influenzae from
TI
                                                                        Page 36
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Haemophilus parainfluenzae.
     Santanam P
ΑU
     EUROPEAN JOURNAL OF CLINICAL MICROBIOLOGY, (1984 Apr) 3 (2) 150-1.
SO
     Journal code: EMY; 8219582. ISSN: 0722-2211.
CY
     GERMANY, WEST: Germany, Federal Republic of
     Letter
DT
     English
LA
     Priority Journals
FS
     198407
EM
     Entered STN: 19900320
ED
     Last Updated on STN: 19900320
     Entered Medline: 19840716
     *Bacteriological Techniques
CT
     *Haemophilus: IP, isolation & purification
     *Haemophilus influenzae: IP, isolation & purification
     *Indicators and Reagents
     *Reagent Strips
                          MEDLINE
    ANSWER 11 OF 15
L18
     83298942
                  MEDLINE
AN
               PubMed ID: 6412201
DN
     An easily prepared and accurate test strip for the detection of
ТΙ
     beta-lactamase production by both gram negative and positive organisms.
ΑU
     Mugg P A
     PATHOLOGY, (1983 Apr) 15 (2) 175-6.
SO
     Journal code: OTA; 0175411. ISSN: 0031-3025.
CY
     Australia
     Journal; Article; (JOURNAL ARTICLE)
DT
LA
     English
     Priority Journals
FS
EM
     198310
     Entered STN: 19900319
ED
     Last Updated on STN: 19900319
     Entered Medline: 19831021
     A simple method is described for the preparation of a paper test
AΒ
     strip for the rapid detection of the beta-lactamase enzymes of
Haemophilus sp., N. gonorrhoeae, S. aureus and S. epidermidis. The test
     strips were compared with the chromogenic cephalosporin technique
     for beta-lactamase detection and found to give 100% agreement. The
     beta-lactamase detection strips were easily prepared, very
     inexpensive, required no special equipment and could be stored for up to
6
     mth.
CT
     Haemophilus: EN, enzymology
     *Indicators and Reagents
      Neisseria gonorrhoeae: EN, enzymology
     *Reagent Strips
      Staphylococcus: EN, enzymology
      Staphylococcus aureus: EN, enzymology
     *beta-Lactamases: AN, analysis
     ANSWER 12 OF 15
                          MEDLINE
L18
     83109707
                   MEDLINE
AN
DN
                PubMed ID: 6337192
     83109707
     Biotyping of Haemophilus using API 10S--an epidemiological tool?.
ΤI
ΑU
     Mehtar S; Afshar S A
     JOURNAL OF CLINICAL PATHOLOGY, (1983 Jan) 36 (1) 96-9.
SO
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Journal code: HT3; 0376601. ISSN: 0021-9746.
CY
     ENGLAND: United Kingdom
DТ
     Journal; Article; (JOURNAL ARTICLE)
LA
     English
     Abridged Index Medicus Journals; Priority Journals
FS
     198303
EM
     Entered STN: 19900318
ED
     Last Updated on STN: 19900318
     Entered Medline: 19830311
     Check Tags: Comparative Study; Human
CT
      Adolescence
      Adult
      Age Factors
      Bacteriological Techniques
      Child
      Child, Preschool
     *Haemophilus: CL, classification
      Haemophilus: IP, isolation & purification
      Haemophilus: ME, metabolism
      Haemophilus Infections: MI, microbiology
      Haemophilus influenzae: CL, classification
Haemophilus influenzae: IP, isolation & purification
      Haemophilus influenzae: ME, metabolism
     *Indicators and Reagents
     *Reagent Strips
     ANSWER 13 OF 15
L18
                         MEDLINE
ΑN
     82167977
                  MEDLINE
     82167977
                PubMed ID: 7040443
DN
     Evaluation of the rapid penicillinase paper strip test for
TТ
     detection of beta-lactamase.
ΑU
     Oberhofer T R; Towle D W
     JOURNAL OF CLINICAL MICROBIOLOGY, (1982 Feb) 15 (2) 196-9.
SO
     Journal code: HSH; 7505564. ISSN: 0095-1137.
CY
     United States
     Journal; Article; (JOURNAL ARTICLE)
DT
LA
     English
FS
     Priority Journals
EM
     198206
     Entered STN: 19900317
ED
     Last Updated on STN: 19900317
     Entered Medline: 19820621
AΒ
     The penicillin-starch paper strip method was compared with the
     acidometric and iodometric methods for assaying beta-lactamase
production,
     using fresh isolates of clinically important bacteria. Results obtained
by
     the three methods were compared for rapidity, accuracy, and stability of
     reagents. Of the 210 isolates tested by the paper strip method,
     301 isolates tested by the acidometric method, and 117 isolates tested by
     the iodometric method, all were in perfect agreement with the disk
     diffusion susceptibility test except one strain each of Haemophilus
     influenzae, Staphylococcus aureus, and Staphylococcus epidermidis. The H.
     influenzae isolate was penicillin resistant and failed to give a positive
     test for beta-lactamase in all three tests. The staphylococci
     (intermediate and resistant in susceptibility, respectively) failed to
     give a positive test for beta-lactamase with the iodometric method. The
                                                                          Page 38
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results of the paper strip method, in which 3,241 strains representing nine species of bacteria were used, correlated completely with disk susceptibility tests except for 2 and 69 strains, respectively, of penicillin-resistant, beta-lactamase-negative H. influenzae and H. parainfluenzae. The results of this study indicate that the paper strip method is accurate, simple to perform, extremely economical, and uses materials that are stable when stored frozen. It is eminently suitable for routine laboratory use.

CT Check Tags: Human; Support, Non-U.S. Gov't

\*Bacteria: EN, enzymology

\*Bacteriological Techniques

Haemophilus: EN, enzymology

Penicillinase: AN, analysis

Reagent Strips

Staphylococcus: EN, enzymology \*beta-Lactamases: AN, analysis

L18 ANSWER 14 OF 15 MEDLINE

AN 77238680 MEDLINE

DN 77238680 PubMed ID: 888594

TI [Polyacrylamid-gel-electrophoresis of Haemophilus proteins (author's transl)].

Elektrophoretische Auftrennung von Haemophilus-Proteinen im Polyacrylamid-Gel.

AU Neumann U; Hinz K H

SO ZENTRALBLATT FUR BAKTERIOLOGIE, PARASITENKUNDE, INFEKTIONSKRANKHEITEN UND HYGIENE. ERSTE ABTEILUNG ORIGINALE. REIHE A: MEDIZINISCHE MIKROBIOLOGIE UND PARASITOLOGIE, (1977 Jun) 238 (2) 244-50.

Journal code: Y52; 0331570. ISSN: 0300-9688.

CY GERMANY, WEST: Germany, Federal Republic of

DT Journal; Article; (JOURNAL ARTICLE)

LA German

FS Priority Journals

EM 197709

ED Entered STN: 19900314 Last Updated on STN: 19900314 Entered Medline: 19770922

After phenol-acetic-acid extraction the following Haemophilus strains resp. their proteins were subjected the polyacrylamid-gel-electrophoresis in presence of 8 M urea: Strains of the serovar A of H. paragallinarum: 0083, 1516, 1598, 2213, 1645, 1646, Lohren, 2671, 1385, 758, 17756; strains of serovar B of H. paragallinarum: 0222, 2600, 733, 2028, 1596, 2026, 1676, 245, the S and R-form of 2403 as well as the strains 782 and 1655, which were not serotyped; strains of H. paravium sp. nova (HINZ: Inst. J. Syst. Bacteriol. in press): 1762, 62 (Serovar 1), 2654, 2659 (Serovar 2), 780 (Serovar 3), 94 (Serovar 4) and 1254, 0002, 0003, which were not serotyped. H. parainfluenzae (NCTC 4101) and H. parasuis were examined in the same way. The Coomassie Blue-stained protein patterns

show

that each of the strains tested developed its characteristic protein pattern, with exception of the S- and R-form of the strain 2403, which developed identical pattern. Interrelations between electrophoretic pattern and biological properties such as biochemical activities or pathogenicity could not be proved. However, the procedure described seems to be suitable for strain- or clon-identification on the subspecies level.

The electrophoresis apparatus, which was made according to our

instructions was less expensive than corresponding available equipments and proved to be usable for the polyacrylamid-gel-electrophoresis. \*Bacterial Proteins: AN, analysis CT\*Electrophoresis, Polyacrylamide Gel \*Haemophilus: AN, analysis Haemophilus: CL, classification Species Specificity L18 ANSWER 15 OF 15 MEDLINE MEDLINE 77110823 ΑN 77110823 PubMed ID: 827984 DN [Qualitative and quantitative researches into bacterial flora of TΤ respiratory apparatus (author's transl)]. Indagini qualitative e quantitative sulla flora batterica dell'apparato respiratorio. Torelli P C; Tortoli E ΑU ANNALI SCLAVO, (1976 Mar-Apr) 18 (2) 198-206. SO Journal code: 65C; 2985177R. ISSN: 0003-472X. CY Italy Journal; Article; (JOURNAL ARTICLE) DTLA Italian Priority Journals FS EM 197703 ED Entered STN: 19900313 Last Updated on STN: 19900313 Entered Medline: 19770331 The first group of data illustrated in this research concern the AB isolation's frequencies of bacterial strains from 7000 samples of respiratory materials examinated in two years, 1973 and 1974, by two different techniques. Significant differences were observed in the isolation's percentage of various bacterial strains; these differences, at least partly, can be attributed at the different techniques employed in two years. The second group of data is represented by results of bacterial quantitation performed in 100 samples of purulent sputum, with respectives clinical notes unknown. Potentially pathogenous strains in pure culture and at a concentration of 10(7) or higher were isolated only in 16 samples of sputum in the other samples normal bacterial flora was isolated alone or associated with potentially pathogenous strains. CT Check Tags: Human Candida albicans: IP, isolation & purification Escherichia coli: IP, isolation & purification Haemophilus: IP, isolation & purification Klebsiella: IP, isolation & purification Neisseria: IP, isolation & purification Pseudomonas aeruginosa: IP, isolation & purification \*Respiratory System: MI, microbiology \*Sputum: MI, microbiology Staphylococcus: IP, isolation & purification

#### => d his

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(FILE 'HOME' ENTERED AT 07:38:54 ON 09 AUG 2001)
     FILE 'BIOSIS' ENTERED AT 07:39:14 ON 09 AUG 2001
            533 S ICT
L1
            307 S IMMUNOCHROMATOGRA?
L2
L3
             31 S L1 AND L2
            809 S L1 OR L2
L4
          18324 S HAEMOPHILUS OR H INFLUENZ?
L5
              1 S L5 AND L4
L6
           3855 S (POLYSACCHARIDE# OR CARBOHYDRATE#) (4A) ANTIGEN#
L7
rs
         487656 S ANTIBOD?
           1745 S L7 AND L8
L9
              0 S L9 AND L4
L10
              1 S L7 AND L4
L11
          50774 S L8 (5A) ANTIGEN#
L12
             18 S L12 AND L4
L13
          36386 S STRIP? OR TESTSTRIP
L14
              4 S L13 AND L14
L15
            597 S ALL 1
L16
=> d all 16;d all 111;d bib ab it 115 1-4
     ANSWER 1 OF 1 BIOSIS COPYRIGHT 2001 BIOSIS
L6
ΑN
     2001:351661 BIOSIS
     PREV200100351661
DN
     Laboratory diagnosis of ocular adenovirus infection: Is there really one
ΤI
     best test.
     Kowalski, R. P. (1); Suzow, J. (1); Karenchak, L. M. (1); Romanowski, E. G. (1); Weck, K. E. (1); Gordon, Y. J. (1)
ΑU
     (1) Charles T. Campbell Ophthalmic Lab and Dept of Molecular Diagnostics,
CS
     Univ of Pittsburgh Med Ctr, Pittsburgh, PA USA
     IOVS, (March 15, 2001) Vol. 42, No. 4, pp. S579. print.
SO
     Meeting Info.: Annual Meeting of the Association for Research in Vision
     and Ophthalmology Fort Lauderdale, Florida, USA April 29-May 04, 2001
DT
     Conference
     English
LA
     English
SL
     Virology - Animal Host Viruses *33506
CC
     General Biology - Symposia, Transactions and Proceedings of Conferences,
     Congresses, Review Annuals *00520
     Biochemical Studies - Nucleic Acids, Purines and Pyrimidines *10062
     Sense Organs, Associated Structures and Functions - Physiology and
     Biochemistry
                   *20004
     Sense Organs, Associated Structures and Functions - Pathology *20006
     Physiology and Biochemistry of Bacteria *31000
     Medical and Clinical Microbiology - Virology *36006
                     02601
     Adenoviridae
     Herpesviridae
                      02612
     Pasteurellaceae
                        06703
     Chlamydiaceae
                      07121
     Micrococcaceae
     Major Concepts
        Infection; Methods and Techniques; Sense Organs (Sensory Reception)
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Page 1

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Parts, Structures, & Systems of Organisms
        eye: sensory system
IT
     Diseases
        ocular adenovirus infection: eye disease, laboratory diagnosis, viral
        disease
    Chemicals & Biochemicals
        DNA: adenoviral
    Methods & Equipment
ΙT
        PCR [polymerase chain reaction]: DNA amplification, genetic method,
        in-situ recombinant gene expression detection, laboratory method,
        sensitivity, sequencing techniques, specificity; enzyme immunoassay
        [Adenoclone]: enzymatic method, laboratory method, sensitivity,
        specificity; immunochromatography [AdenoTest]: laboratory
        method, sensitivity, specificity; shell vial culture: culture method,
        laboratory method, sensitivity, specificity
IT
     Miscellaneous Descriptors
        processing time; Meeting Abstract
ORGN Super Taxa
        Adenoviridae: Animal Viruses, Viruses, Microorganisms; Animalia;
        Chlamydiaceae: Chlamydiales, Rickettsias and Chlamydias, Eubacteria,
        Bacteria, Microorganisms; Herpesviridae: Animal Viruses, Viruses,
        Microorganisms; Micrococcaceae: Gram-Positive Cocci, Eubacteria,
        Bacteria, Microorganisms; Pasteurellaceae: Facultatively Anaerobic
        Gram-Negative Rods, Eubacteria, Bacteria, Microorganisms
ORGN Organism Name
        Chlamydia (Chlamydiaceae); HSV [Herpes Simplex Virus] (Herpesviridae);
     Haemophilus (Pasteurellaceae); S. aureus [Staphylococcus
        aureus] (Micrococcaceae); St. pneumo [Staphylococcus pneumoniae]
        (Micrococcaceae); adenovirus (Adenoviridae): pathogen; animal
        (Animalia): host
ORGN Organism Superterms
        Animal Viruses; Animals; Bacteria; Eubacteria; Microorganisms; Viruses
    ANSWER 1 OF 1 BIOSIS COPYRIGHT 2001 BIOSIS
L11
     2001:133082 BIOSIS
AN
     PREV200100133082
DN
     Detection of Streptococcus pneumoniae antigen by a rapid
ΤI
     immunochromatographic assay in urine samples.
     Dominguez, Jose (1); Gali, Nuria; Blanco, Silvia; Pedroso, Pablo; Prat, Cristina; Matas, Lurdes; Ausina, Vicente
ΑU
     (1) Servei de Microbiologia, Hospital Universitari Germans Trias i Pujol,
CS
     Carretera del Canyet s/n, 08916, Badalona, Catalonia:
     jadoming@ns.hugtip.scs.es Spain
     Chest, (January, 2001) Vol. 119, No. 1, pp. 243-249. print.
SO
     ISSN: 0012-3692.
DT
     Article
LΑ
     English
SL
     English
     Study objectives: Evaluation of a newly available rapid (15 min)
AB
     immunochromatographic membrane test (ICT) to detect
     Streptococcus pneumoniae in urine samples, in order to assess its utility
     in the diagnosis of bacteremic and nonbacteremic pneumococcal pneumonia.
     Design: Retrospective study. Setting: We studied urine samples from 51
     patients with bacteremic and nonbacteremic pneumonia due to S pneumoniae
                                                                          Page 2
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diagnosed by blood culture and pneumococcal polysaccharide capsular antigen detection by counterimmunoelectrophoresis in urine samples, 16 patients with probable pneumococcal pneumonia, 71 patients with nonpneumococcal pneumonia, and 16 patients with pneumonia but no pathogen identified. Urine samples were collected and frozen at -20degreeC until used. The ICT test was performed following the instructions of the manufacturer. Measurements and results: S pneumoniae antigen was detected in 41 of 51 patients with pneumococcal pneumonia (80.4%); results were positive in 23 of 28 bacteremic cases (82.1%) and 18 of 23 nonbacteremic cases (78.3%). From patients with a diagnosis of presumptive pneumococcal pneumonia, antigen was detected in seven urine samples (43.7%) and also in one case of the 16 patients with pneumonia

but

no pathogen identified. The specificity of the ICT test was 97.2%. Conclusion: The ICT assay is a valuable tool for the diagnosis of pneumococcal pneumonia, especially for the nonbacteremic cases.

Respiratory System - Pathology \*16006 Clinical Biochemistry; General Methods and Applications \*10006 Physiology and Biochemistry of Bacteria \*31000 Medical and Clinical Microbiology - Bacteriology \*36002

Gram-Positive Cocci 07700 BC

ΙT Major Concepts Clinical Chemistry (Allied Medical Sciences); Infection; Pulmonary Medicine (Human Medicine, Medical Sciences)

ΙT non-pneumococcal pneumonia: respiratory system disease; pneumococcal pneumonia: bacterial disease, respiratory system disease

Chemicals & Biochemicals ΙT

Streptococcus pneumoniae antigen

Alternate Indexing ΙT

Pneumonia, Pneumococcal (MeSH)

Methods & Equipment IT

immunochromatographic membrane test: detection method, serodiagnostic method

ORGN Super Taxa

in

Gram-Positive Cocci: Eubacteria, Bacteria, Microorganisms; Hominidae: Primates, Mammalia, Vertebrata, Chordata, Animalia

ORGN Organism Name

Streptococcus pneumoniae (Gram-Positive Cocci): pathogen; human (Hominidae): patient

ORGN Organism Superterms

Animals; Bacteria; Chordates; Eubacteria; Humans; Mammals; Microorganisms; Primates; Vertebrates

- ANSWER 1 OF 4 BIOSIS COPYRIGHT 2001 BIOSIS L15
- 2000:497615 BIOSIS AN
- PREV200000497736 DN
- Development of rapid one-step immunochromatographic assay. ΤI
- Paek, Se-Hwan (1); Lee, Seung-Hwa; Cho, Joung-Hwan; Kim, Young-Sang ΑU
- (1) Graduate School of Biotechnology, Korea University, 5-ka, Anam-dong, CS Sungbuk-ku, Seoul, 136-701 South Korea
- Methods (Orlando), (September, 2000) Vol. 22, No. 1, pp. 53-60. print. SO

ISSN: 1046-2023. Article DT English LA SLEnglish An analytical system for a one-step immunoassay has been constructed AΒ using the concept of immunochromatography. The system employed two different antibodies that bound distinct epitopes of an analyte molecule: an antibody labeled with a signal generator (e.g., colloidal gold), which was placed in the dry state at a predetermined site on a glass-fiber membrane, and another antibody immobilized on the surface of a nitrocellulose membrane. Three membranes, one with the tracer, one with immobilized antibody, and a cellulose membrane as the absorbent of medium (in a sequence from the bottom), were attached to a plastic film and cut into strips. Aqueous medium containing analyte absorbed from the bottom end of the immunostrip dissolved the labeled antibody, and the antigen-antibody binding complex formed was transported into the next nitrocellulose membrane by the flow caused by capillary action. The complex subsequently reacted with the immobilized antibody, which generated a signal in proportion to the analyte concentration. The convective mass transfer of the immunoreactant to the binding partner allowed the assay to be performed with no handling of reagents. The reaction, however, was carried out under nonequilibrium conditions, which resulted in decreased sensitivity as compared with assays performed in an equilibrium mode (e.g., ELISA). To minimize such sacrifice, major factors that control system performance were identified and the system was then devised under optimal conditions. ΙT Major Concepts Methods and Techniques Chemicals & Biochemicals ፐጥ antibodies ΙT Methods & Equipment ELISA: analytical method, detection/labeling techniques; immunochromatography: analytical method, chromatographic techniques; nitrocellulose membrane: Millipore, equipment ANSWER 2 OF 4 BIOSIS COPYRIGHT 2001 BIOSIS L15 1998:367509 BIOSIS ANDN PREV199800367509 Rapid accurate field diagnosis of Indian visceral leishmaniasis. ΤI Sundar, Shyam; Reed, Steven G.; Singh, Vijay P.; Kumar, Prasanna C. K.; ΑU Murray, Henry W. (1) (1) Box 130, 535 E. 68th St., New York, NY 10021 USA CS Lancet (North American Edition), (Feb. 21, 1998) Vol. 351, No. 9102, pp. SO 563-565. ISSN: 0099-5355. DT Article LA English Background: A firm diagnosis of visceral leishmaniasis (kala-azar) requires demonstration of the parasite in organ aspirates or tissue samples. The aim of this prospective study was to assess the diagnostic usefulness of non-invasive testing for antibody to the leishmanial antigen K39 by means of antigen-impregnated nitrocellulose paper strips adapted for use under field conditions. Methods: One drop of peripheral blood is applied to the

nitrocellulose strip, Three drops of test buffer

(phosphate-buffered saline plus bovine serum albumin) are added to the dried blood. The development of two visible bands indicates presence of IgG anti-K39. 323 consecutive patients with suspected kala-azar referred to two specialist units in India, and 25 healthy controls, provided fingerstick blood samples for the test. Spleen aspirates were taken from 250 patients. Findings: Kala-azar was confirmed by microscopy of spleen-aspirate smears in 127 patients. The K39 **strip** test was positive in all 127; the estimated sensitivity was therefore 100% (95%)

CI

98-100). Four patients had positive strip tests but negative aspirate smears; all four responded to treatment for leishmaniasis. 217 individuals, including the 25 healthy controls, 73 patients with malaria or tuberculosis, and 119 spleen-aspirate-negative patients who had presumed malaria or cirrhosis (79) or no final diagnosis (40), had negative strip-test results. None of the 119 aspirate-negative patients developed evidence of kala-azar during 3-6 months of follow-up. The estimated specificity of the strip test was 98% (95-100; 217/221). Interpretation: Detection of anti-K39 by immunochromatographic strip testing is a rapid and non-invasive method of diagnosing kala-azar, which has good sensitivity and specificity and is well suited for use in field conditions.

IT Major Concepts
Clinical Immunology (Human Medicine, Medical Sciences); Methods and
Techniques; Parasitology

IT Diseases

visceral leishmaniasis [kala-azar]: integumentary system disease, parasitic disease

IT Chemicals & Biochemicals

immunoglobulin G; recombinant K39 antigen

IT Methods & Equipment

anti-K39 antigen detection: diagnostic method, immunological method; immunochromatographic strip test: diagnostic method,

field method, rapid, immunological method

GT India (Oriental region)

ORGN Super Taxa

Flagellata: Protozoa, Invertebrata, Animalia; Hominidae: Primates, Mammalia, Vertebrata, Chordata, Animalia

ORGN Organism Name

human (Hominidae): parasite host, patient; Leishmania-chagasi (Flagellata): parasite; Leishmania-donovani (Flagellata): parasite; Leishmania-infantum (Flagellata): parasite

ORGN Organism Superterms

Animals; Chordates; Humans; Invertebrates; Mammals; Microorganisms; Primates; Protozoans; Vertebrates

- L15 ANSWER 3 OF 4 BIOSIS COPYRIGHT 2001 BIOSIS
- AN 1996:285319 BIOSIS
- DN PREV199699007675
- TI Evaluation of immunochromatographic assay systems for rapid detection of hepatitis B surface antigen and antibody, Dainascreen HBsAg and Dainascreen Ausab.
- AU Sato, Kumiko; Ichiyama, Satoshi (1); Iinuma, Yoshitsugu; Nada, Toshi; Shimokata, Kaoru; Nakashima, Nobuo
- CS (1) Dep. Clin. Lab. Med., Nagoya Univ. Hosp., 65 Tsurumai-cho, Showa-ku, Nagoya 466 Japan
- SO Journal of Clinical Microbiology, (1996) Vol. 34, No. 6, pp. 1420-1422. ISSN: 0095-1137.

```
Article
DT
    English
LA
    We evaluated two immunochromatographic assays (ICAs),
AB
     Dainascreen HBsAq for detecting human hepatitis B surface antigen (HBsAg)
     and Dainascreen Ausab for detecting human hepatitis B surface antibody
     (anti-HBs) in human serum. The ICA systems are composed of a comb-shaped
     device that contains nitrocellulose strips on which complexes of
     HBsAg and anti-HBs can be visualized. The results can be read within 15
    min of incubation. The limit of detection for HBsAg was 3.1 ng/ml, and
     that for anti-HBs was 42 mIU/ml. Results of HBsAg detection agreed
     completely with those of conventional enzyme immunoassays (EIAs) and
     showed a 100% sensitivity (158 of 158 samples) and a 100% specificity
(304
     of 304 samples). The Dainascreen Ausab detected 184 of the 199
     EIA-positive samples (sensitivity, 92.5%) and yielded 6 positive results
     among the 281 EIA-negative samples (specificity, 97.9%). The ICA systems
     are rapid and sensitive methods for detecting HBsAg and anti-HBs. They
are
     low-cost systems that need no complex instrumentation for analysis and
can
    be recommended for routine use in clinical microbiology laboratories.
    Major Concepts
TT
        Immune System (Chemical Coordination and Homeostasis); Infection;
        Pathology; Serology (Allied Medical Sciences)
    Miscellaneous Descriptors
IT
        DIAGNOSTIC METHOD; HUMAN SERUM
ORGN Super Taxa
        Hepadnaviridae: Viruses; Hominidae: Primates, Mammalia, Vertebrata,
        Chordata, Animalia
ORGN Organism Name
        Hepadnaviridae (Hepadnaviridae); Hominidae (Hominidae)
ORGN Organism Superterms
        animals; chordates; humans; mammals; microorganisms; primates;
        vertebrates; viruses
    ANSWER 4 OF 4 BIOSIS COPYRIGHT 2001 BIOSIS
L15
     1985:162782 BIOSIS
ΑN
DN
     BR29:52778
     STRIP ASSAY FOR QUANTITATION OF ANTIBODY TO HEPATITIS
ΤI
     B SURFACE ANTIGEN.
     VAN HAMONT J E; VINCENT J W; PAPPAS M G; HAJKOWSKI R; SETTERSTROM J A
ΑU
     U.S. ARMY INSTITUTE OF DENTAL RESEARCH, WASHINGTON, D.C.
CS
     85TH ANNUAL MEETING OF THE AMERICAN SOCIETY FOR MICROBIOLOGY, LAS VEGAS,
SO
     NEV., USA, MAR. 3-7, 1985. ABSTR ANNU MEET AM SOC MICROBIOL. (1985) 85
     (0), 390.
     CODEN: ASMACK. ISSN: 0094-8519.
     Conference
DT
     BR; OLD
FS
LΑ
     English
IT
     Miscellaneous Descriptors
        ABSTRACT HUMAN MOUSE RADIOIMMUNOASSAY IMMUNOCHROMATOGRAPHY
```

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=> d his
     (FILE 'BIOSIS' ENTERED AT 07:39:14 ON 09 AUG 2001)
                DEL HIS Y
     FILE 'STNGUIDE' ENTERED AT 07:48:38 ON 09 AUG 2001
     FILE 'WPIDS' ENTERED AT 07:54:13 ON 09 AUG 2001
            178 S IMMUNOCHROMATOGRA? OR ICT OR IMMUNO CHROMATOGRAP?
L1
                  (CARBOHYDRATE? OR POLYSACCAHARIDE# OR SACCHARIDE#)
L2
          28904 S L2 OR POLYSACCHARIDE#
L3
L4
            462 S L3 (5A) ANTIGEN#
              2 S L4 AND L1
L5
=> d .wp 1-2
     ANSWER 1 OF 2 WPIDS COPYRIGHT 2001 DERWENT INFORMATION LTD
L5
                        WPIDS
     2000-283451 [24]
ΑN
DNC
    C2000-085570
     Obtaining cell wall C-polysaccharide antigens
ΤI
     containing not more than 10% protein from the bacterium Streptococcus
     pneumoniae for the production of antigen-specific antibodies.
DC
     B04 D16
     FENT, M K; KOULCHIN, V A; MOLOKOVA, E V; MOORE, N J
ΙN
     (BINA-N) BINAX INC
PΑ
CYC
    36
     WO 2000016803 A1 20000330 (200024)* EN
                                              q8E
PΙ
        RW: AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE
         W: AT AU CA CH CN CZ DE DK ES FI GB HU IL IN JP KR LU MX NO NZ PL PT
            RU SE SK UA ZA
     AU 9961513
                   A 20000410 (200035)
                   A1 20010711 (200140)
     EP 1113817
                                         EN
         R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE
    WO 2000016803 A1 WO 1999-US21505 19990920; AU 9961513 A AU 1999-61513
ADT
     19990920; EP 1113817 A1 EP 1999-948305 19990920, WO 1999-US21505 19990920
    AU 9961513 A Based on WO 200016803; EP 1113817 A1 Based on WO 200016803
FDT
                      19990916; US 1998-156486
                                                 19980918
PRAI US 1999-397110
     WO 200016803 A UPAB: 20000522
     NOVELTY - Obtaining a cell wall C-polysaccharide antigen
     containing not more than 10% protein from the bacterium Streptococcus
     pneumoniae for the production of antigen-specific antibodies, is new.
          DETAILED DESCRIPTION - Obtaining a cell wall C-polysaccharide
     antigen containing not more than 10% protein from the bacterium
     Streptococcus pneumoniae comprises:
          (a) culturing the bacterium for a time requisite to obtain a sample
     of desired size and harvesting the bacterial cells in the form of a wet
     pellet;
          (b) suspending the wet cell pellet in an alkaline solution and
     mixing;
          (c) adjusting the pH to an acid pH with a strong acid and
     centrifuging;
          (d) separating a supernatant and adjusting its pH to approximate
```

(e) digesting this product with a broad spectrum protease enzyme

Page 7

neutrality;

preparation to destroy residual proteins;

- (f) adjusting the pH to the alkaline side with a weakly alkaline aqueous solution;
- (g) separating out the essentially protein free carbohydrate or polysaccharide antigen on a size exclusion column equilibrated with a weakly alkaline solution; and
  - (h) pooling material eluted in the first peak and adjusting its pH

to

approximate neutrality.

INDEPENDENT CLAIMS are also included for the following:

- (1) a cell wall C-polysaccharide antigen obtained from the method above;
- (2) a method for purifying raw antibodies to S. pneumoniae comprising:
- (a) separating from S. pneumoniae bacteria a cell wall C-polysaccharide antigen containing not more than 10% protein;
- (b) conjugating the antigen to one end of a two-ended spacer molecule;
  - (c) coupling the conjugate to an activated chromatographic column;
- (d) subjecting the raw antibodies to affinity chromatography on the column from (c) to obtain purified antigen-specific antibodies; and
- (e) eluting from the column the purified antigen-specific antibodies;
  - (3) purified antigen-specific antibodies obtained by (2);
  - (4) a chromatographic column for affinity purification of raw antibodies to S. pneumoniae having, via a spacer molecule, a coupled purified C-polysaccharide cell wall antigen of S. pneumoniae containing not more than 10% protein;
  - (5) a method of assaying for the presence of S. pneumoniae or its cell wall C-polysaccharide antigen in a fluid comprising:
- (a) extracting from a culture of S. pneumoniae bacteria the cell wall

C-polysaccharide antigen containing not more than 10% protein;

- (b) coupling the antigen to a spacer molecule to form a conjugate;
- (c) coupling the conjugate to a chromatographic affinity column;
- (d) purifying raw antibodies to S. pneumoniae bacteria with the chromatographic affinity column of (c) to produce purified antigen-specific antibodies; and
- (e) using the purified antibodies of (d) to detect the presence/absence of S. pneumoniae or its C-polysaccharide cell wall antigen in a fluid;
- (6) an immunochromatographic (ICT) assay for the detection of S. pneumoniae bacteria or the C-polysaccharide cell wall antigen of the bacteria comprising:
- (a) contacting a sample of a fluid suspected of containing the bacteria or their antigen with an ICT device comprising a strip of a bibulous material comprising:
- (i) a first zone in which has been embedded a conjugate of a labeling agent that displays a visible color change upon reaction of antibodies

with their corresponding antigenic binding partner and purified antigen-specific antibodies to the C-polysaccharide cell wall antigen of S. pneumoniae, the antibodies having been purified by passage over a chromatographic affinity column to which is

conjugated a purified C-polysaccharide cell wall antigen of S. pneumoniae containing not more than 10% protein; and

- (ii) a second zone having bound the same purified antigen-specific antibodies in unconjugated form, which zone is equipped with a window for viewing color changes;
  - (b) allowing the sample to flow laterally along the test strip to

the

first zone;

- (c) allowing the sample, together with the conjugate of antigen-specific antibodies and label, to flow laterally along the test strip to the second zone; and
- (d) within 15 minutes from the commencement of step (a) observing through the window whether a line of color has appeared in the second zone, indicating the presence in the sample of S. pneumoniae or its cell wall C-polysaccharide antigen; and
  - (7) an ICT device used in (6).
- USE The method is useful for generating antigens which, when conjugated to a spacer molecule, may be attached to a chromatographic affinity column. The column is then used to purify raw antibodies to Streptococcus pneumoniae. The immunochromatographic assay is used for rapid and reliable diagnosis of pathogenic states caused by S. pneumoniae including pneumonia, bronchitis, otitis media, sinusitis and meningitis. Dwg.0/4
- L5 ANSWER 2 OF 2 WPIDS COPYRIGHT 2001 DERWENT INFORMATION LTD
- AN 2000-237549 [20] WPIDS
- DNC C2000-072236
- TI Obtaining protein-free carbohydrate or polysaccharide antigen from a bacterium useful for immunological assays for the detection of Legionella caused diseases.
- DC B04 D16
- IN KOULCHIN, V A; MOLOKOVA, E V; MOORE, N J
- PA (BINA-N) BINAX INC
- CYC 36
- PI WO 2000010584 A1 20000302 (200020)\* EN 31p
  - RW: AT BE CH CY DE DK ES FI FR GB GR IE IT LU MC NL PT SE
  - W: AT AU CA CH CN CZ DE DK ES FI GB HU IL IN JP KR LU MX NO NZ PL PT RU SE SK UA ZA
  - AU 9956933 A 20000314 (200031)
  - EP 1107773 A1 20010620 (200135) EN
  - R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE
- ADT WO 2000010584 A1 WO 1999-US19506 19990825; AU 9956933 A AU 1999-56933 19990825; EP 1107773 A1 EP 1999-943941 19990825, WO 1999-US19506 19990825
- FDT AU 9956933 A Based on WO 200010584; EP 1107773 Al Based on WO 200010584
- PRAI US 1998-139720 19980825
- AB WO 200010584 A UPAB: 20000426
  - NOVELTY A method (I) for obtaining a protein free carbohydrate
  - or polysaccharide antigen from a known Legionella
  - bacterium or serogroup of species, is new.
  - DETAILED DESCRIPTION A method (I) for obtaining a protein free carbohydrate or polysaccharide antigen from a
  - known Legionella bacterium or serogroup of species, is new and comprises:
  - (1) culturing the bacterium to obtain a sample of desired size and harvesting the bacterial cells in the form of a wet cell pellet;
  - (2) suspending the wet cell pellet in an alkaline solution and mixing;
    - (3) adjusting the pH to an acid pH with a strong acid and

centrifuging;

- (4) separating the supernatant and adjusting its pH to approximate neutrality;
- (5) digesting the product with a broad spectrum protease enzyme preparation to destroy residual proteins;
  - (6) adjusting the pH to alkaline with a weak alkaline solution; and
  - (7) pooling material eluted in the first peak and adjusting its pH

to

approximate neutrality.

INDEPENDENT CLAIMS are also included for the following:

- (1) a protein free carbohydrate or polysaccharide antigen of a bacterium obtained from a known Legionella pneumophila species or serogroup of species using (I);
- (2) a method (II) for the purification of raw antibodies to a species

or serogroup of a species of Legionella bacteria, comprising:

- (a) separating from the same species or serogroup of a species of Legionella bacteria, a protein free carbohydrate or polysaccharide antigen;
- (b) conjugating the antigen to one end of the a two ended spacer molecule to form a conjugate of the protein free antigen with the spacer molecule;
  - (c) coupling the conjugate to an activated chromatographic column;
- (d) subjecting the raw antibodies to affinity chromatography on the column from (c) to obtain purified antigen specific antibodies; and
  - (e) eluting from the column the purified antigen-specific

antibodies;

- (3) purified antigen-specific antibodies to a bacterium of at least one species or serogroup of species of L. pneumophila obtained by (II);
- (4) a chromatographic column (III) for affinity purification of raw antibodies in method (II);
- (5) a method (IV) for assaying for the presence of Legionella bacteria or their antigenic components in a fluid comprising purifying

raw

antibodies specific to L. pneumophila species and using the purified antibodies of to detect the presence or absence of the corresponding Legionella bacteria or their antigens in a fluid;

(6) a process (V) for detecting the presence of at least one species or serogroup of a species of Legionella bacteria or its antigen in a fluid

medium, where the detecting agent is antigen specific Legionella antibodies obtained by (II); and

- (7) an ICT (immunochromatographic test) assay
- (VI) for the detection of at least one species or serogroup of a species of Legionella bacteria or antigens of the bacteria, comprising:
- (a) contacting a sample of a fluid suspected of containing the bacteria or their antigen with an ICT device comprising a strip of bibulous material comprising:
- (i) a zone with a conjugate of a labelling agent (A) embedded in it, that displays a visible color change upon reaction of antibodies with their corresponding antigenic binding partner and purified antigen-specific conjugated antibodies (B) to the Legionella species to

be

detected; and

(ii) a second zone having bound the same purified antigen specific antibodies in unconjugated form which is equipped with a window for viewing color changes;

- (b) allowing the sample to flow laterally along the test strip to the first zone;
  - (c) allowing the sample together with the conjugate of affinity purified antibodies and label to flow laterally along the test strip to the second zone and;
  - (d) within approximately 15 minutes from the commencement of (a), observing through the window whether a line of color has appeared in the second zone thereby indicating the presence in the sample of the Legionella bacteria species or serogroup that is sought to be detected.
  - USE The purified antigens are useful for the affinity purification of polyvalent antibodies to corresponding Legionella organisms. The methods are also useful for the detection of Legionella caused diseases such as Legionnaires disease and Pontiac fever in humans and for the detection of environmental sources of Legionella infectious agents.

    ADVANTAGE The ICT has the ability to give a test result within a
- minute time span for the presence or absence of L. pneumophila serogroup  ${\bf 1}$ 
  - (or its antigen) which results in high specificity and sensitivity. The test can be reliably and quickly conducted to yield a result of high specificity and high sensitivity and this is believed to be due to the strongly reactive nature of the affinity purified antibodies. Dwg.0/2

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=> d his
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(FILE 'MEDLINE' ENTERED AT 07:58:03 ON 09 AUG 2001)
                DEL HIS Y
                E ANTIGENS, BACTERIAL+NT/CT
          68803 S ANTIGENS, BACTERIAL+NT/CT
L1
            884 S ICT OR IMMUNOCHROMATOGRA?
L2
L3
            19 S L1 AND L2
                E HAEMOPHILUS/CT
                                         includes all Haemophilus species
                E E3+ALL
          15898 S PASTEURELLACEAE+NT/CT
L4
              0 S L4 AND L3
L5
=> d .med 1-19 13
    ANSWER 1 OF 19
                       MEDLINE
L3
     2001382101 MEDLINE
ΑN
    21145371 PubMed ID: 11248521
DN
    On-site diagnosis of H. pylori infection by urine.
TΙ
    Miwa H; Akamatsu S; Tachikawa T; Sogabe T; Ohtaka K; Nagahara A; Sugiyama
ΑU
     Y; Sato N
     Department of Gastroenterology, Juntendo University School of Medicine,
CS
     2-1-1 Hongo, Bunkyoku, 113-8421, Tokyo, Japan...
miwahqi@med.juntendo.ac.jp
    DIAGNOSTIC MICROBIOLOGY AND INFECTIOUS DISEASE, (2001 Feb) 39 (2) 95-7.
     Journal code: DMI; 8305899. ISSN: 0732-8893.
    United States
CY
     (EVALUATION STUDIES)
DT
     Journal; Article; (JOURNAL ARTICLE)
LA
    English
FS
    Priority Journals
EM
     200107
    Entered STN: 20010709
ΕD
    Last Updated on STN: 20010709
    Entered Medline: 20010705
AB
    We have recently developed an on-site diagnostic kit for H. pylori
     infection using urine (utilizing immunochromatographic method
     employing a nitrocellulose membrane coated by extracted H. pylori
     antigen). Accordingly, we investigated its usefulness in 155 consecutive
    dyspeptic patients using the 13C urea breath test as a gold standard and
     further compared its performance with two commercially available rapid
    diagnostic kits that use whole blood (Helisal Rapid Blood, and ImmunoCard
    H. pylori). As the results, the urine based on-site diagnostic kit
     provided 95.9% sensitivity and 87.9% specificity with 92.9% accuracy,
    which were comparable or even better than that of both rapid whole blood
    tests, suggesting its usefulness in screening of H. pylori infection.
CT
    Check Tags: Human
     Antigens, Bacterial: BL, blood
     Breath Tests
     *Helicobacter Infections: DI, diagnosis
     *Helicobacter Infections: MI, microbiology
     *Helicobacter pylori: IP, isolation & purification
      Reagent Kits, Diagnostic
      Sensitivity and Specificity
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Urea: ME, metabolism

\*Urine: MI, microbiology ANSWER 2 OF 19 MEDLINE 1.3 AN · 2001323171 MEDLINE PubMed ID: 11238230 21137968 DN Early diagnosis of scrub typhus with a rapid flow assay using recombinant ΤI major outer membrane protein antigen (r56) of Orientia tsutsugamushi. Ching W M; Rowland D; Zhang Z; Bourgeois A L; Kelly D; Dasch G A; Devine ΑU Р Viral and Ricksettial Diseases Department, Infectious Diseases CS Directorate, Code 41, Naval Medical Research Center, Bethesda, Maryland, USA.. Chingw@nmrc.navy.mil CLINICAL AND DIAGNOSTIC LABORATORY IMMUNOLOGY, (2001 Mar) 8 (2) 409-14. SO Journal code: CB7; 9421292. ISSN: 1071-412X. CY United States Journal; Article; (JOURNAL ARTICLE) DΤ LA English FS Priority Journals EM200106 Entered STN: 20010611 F.D Last Updated on STN: 20010611 Entered Medline: 20010607 The variable 56-kDa major outer membrane protein of Orientia AΒ tsutsugamushi is the immunodominant antigen in human scrub typhus infections. We developed a rapid immunochromatographic flow assay (RFA) for the detection of immunoglobulin M (IgM) and IgG antibodies to 0. tsutsugamushi. The RFA employs a truncated recombinant 56-kDa protein from the Karp strain as the antigen. The performance of the RFA was evaluated with a panel of 321 sera (serial bleedings of 85 individuals suspected of scrub typhus) which were collected in the Pescadore Islands, Taiwan, from 1976 to 1977. Among these 85 individuals, IgM tests were negative for 7cases by both RFA and indirect fluorescence assay (IFA) using Karp whole-cell antigen. In 29 cases specific responses were detected by the RFA earlier than by IFA, 44 cases had the same detection time, and 5 cases were detected earlier by IFA than by RFA. For IgG responses, 4 individuals were negative with both methods, 37 cases exhibited earlier detection by RFA than IFA, 42 cases were detected at the same time, and 2 cases were detected earlier by IFA than by RFA. The sensitivities of RFA detection o f antibody in sera from confirmed cases were 74 and 86% for IgM and IgG, respectively. When IgM and IgG results were combined, the sensitivity was 89%. A panel of 78 individual sera collected from patients with no evidence of scrub typhus was used to evaluate the specificity of the RFA. The specificities of the RFA were 99% for IgM and 97% for IgG. The sensitivities of IFA were 53 and 73% for IgM and IgG, respectively, and were 78% when the results of IgM and IgG were combined. The RFA test was significantly better than the IFA test for the early detection of antibody to scrub typhus in primary infections, while both tests were equally sensitive with reinfected individuals. Check Tags: Human; Support, U.S. Gov't, Non-P.H.S.

Page 13

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Antibody Specificity
      Antigens, Bacterial: IM, immunology
     *Bacterial Outer Membrane Proteins: IM, immunology
      IgG: BL, blood
      IqM: BL, blood
     *Immunoassay: MT, methods
      Immunodominant Epitopes: IM, immunology
      Orientia tsutsugamushi: IM, immunology
     *Orientia tsutsugamushi: IP, isolation & purification
     *Reagent Strips
      Recombinant Proteins: IM, immunology
     *Scrub Typhus: DI, diagnosis
      Sensitivity and Specificity
                        MEDLINE
L3
    ANSWER 3 OF 19
AN
     2001318805
                    MEDLINE
               PubMed ID: 11393288
DN
     21284474
     Detection of Legionella pneumophila antigen in urine samples by the
     BinaxNOW immunochromatographic assay and comparison with both
     Binax Legionella Urinary Enzyme Immunoassay (EIA) and Biotest Legionella
     Urin Antigen EIA.
     Helbig J H; Uldum S A; Luck P C; Harrison T G
ΑU
     Medical Microbiology and Hygiene Institute, Technical University Dresden,
CS
     Germany.. Juergen. Helbig@mailbox.tu-dresden.de
SO
     JOURNAL OF MEDICAL MICROBIOLOGY, (2001 Jun) 50 (6) 509-16.
     Journal code: J2N; 0224131. ISSN: 0022-2615.
CY
     England: United Kingdom
     Journal; Article; (JOURNAL ARTICLE)
DT
LA
     English
     Priority Journals
FS
EM
     200106
     Entered STN: 20010625
ΕD
     Last Updated on STN: 20010625
     Entered Medline: 20010621
     The new BinaxNOW Immunochromatographic (ICT) Assay for
AB
     the detection of Legionella pneumophila antigens was used to test 535
     urine specimens from patients with and without Legionnaires' disease. The
     specificity, calculated by testing 112 samples from patients with
     pneumonia of aetiologies other than Legionella infection, and 167 urine
    specimens from urinary tract infections, was found to be 97.1% if the manufacturer's guidelines were followed. However, it was determined that
     the 'false positive' results characterised by very weak bands could be
     discounted by re-examination of the results at 60 min, yielding a
     specificity of 100%. With this minor modification of the procedure
applied
     to examination of urine samples from 117 patients with legionellosis
     confirmed by isolation of L. pneumophila and 70 patients who had
     seroconverted to L. pneumophila serogroup 1, sensitivity was calculated
10
     be 79.7%. In comparison, the sensitivities of the Binax Urinary Antigen
     Enzyme Immunoassay (EIA) and Biotest Urin Antigen EIA were estimated to
be
     79.1 and 83.4%, respectively. Eleven cases (5.9%) were positive by
     BinaxNOW assay but negative by Binax or Biotest EIA, or both. The
     sensitivities of all assays increased to c. 94% if only diagnosis of
cases
     confirmed by isolation of serogroup 1 L. pneumophila was considered,
                                                                          Page 14
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although the sensitivity for infections caused by L. pneumophila serogroup 1 monoclonal antibody (MAb) subgroup Bellingham was significantly lower than for other MAb subgroups. The Biotest EIA recognised 10 (45%) of the 22 cases not caused by L. pneumophila serogroup 1, whereas the two Binax kits detected only three each. The ICT assay BinaxNOW can be recommended as a rapid specific test for the diagnosis of Legionnaires' diseases caused by L. pneumophila serogroup 1, although very weak bands should be interpreted cautiously. Check Tags: Comparative Study; Human CTAntibodies, Monoclonal: IM, immunology \*Antigens, Bacterial: UR, urine Chromatography: MT, methods False Positive Reactions \*Immunoenzyme Techniques: MT, methods Legionella pneumophila: CL, classification \*Legionella pneumophila: IM, immunology Legionella pneumophila: IP, isolation & purification \*Legionnaires' Disease: DI, diagnosis Legionnaires' Disease: UR, urine Reagent Kits, Diagnostic Sensitivity and Specificity Serotyping Time Factors ANSWER 4 OF 19 MEDLINE L3 AN 2001155887 MEDLINE DN 21097861 PubMed ID: 11157611 Detection of Streptococcus pneumoniae antigen by a rapid ΤI immunochromatographic assay in urine samples. Comment in: Chest. 2001 Jan; 119(1):9-11 ·CM Dominguez J; Gali N; Blanco S; Pedroso P; Prat C; Matas L; Ausina V ΑU CS Servei de Microbiologia, Hospital Universitari Germans Trias i Pujol, Badalona, Facultat de Medicina de la Universitat Autonoma de Barcelona, Spain.. jadoming@ns.hugtip.scs.es CHEST, (2001 Jan) 119 (1) 243-9. SO Journal code: D1C; 0231335. ISSN: 0012-3692. United States CY DTJournal; Article; (JOURNAL ARTICLE) LA English FS Abridged Index Medicus Journals; Priority Journals EM 200103 ED Entered STN: 20010404 Last Updated on STN: 20010404 Entered Medline: 20010322 STUDY OBJECTIVES: Evaluation of a newly available rapid (15 min) AB immunochromatographic membrane test (ICT) to detect Streptococcus pneumoniae in urine samples, in order to assess its utility in the diagnosis of bacteremic and nonbacteremic pneumococcal pneumonia. DESIGN: Retrospective study. SETTING: We studied urine samples from 51 patients with bacteremic and nonbacteremic pneumonia due to S pneumoniae diagnosed by blood culture and pneumococcal polysaccharide capsular antigen detection by counterimmunoelectrophoresis in urine samples, 16 patients with probable pneumococcal pneumonia, 71 patients with nonpneumococcal pneumonia, and 16 patients with pneumonia but no pathogen identified. Urine samples were collected and frozen at - 20 degrees C until used. The ICT test was performed following the Page 15

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instructions of the manufacturer. MEASUREMENTS AND RESULTS: S. pneumoniae
     antigen was detected in 41 of 51 patients with pneumococcal pneumonia
     (80.4%); results were positive in 23 of 28 bacteremic cases (82.1%) and
in
     18 of 23 nonbacteremic cases (78.3%). From patients with a diagnosis of
     presumptive pneumococcal pneumonia, antigen was detected in seven urine
     samples (43.7%) and also in one case of the 16 patients with pneumonia
but
     no pathogen identified. The specificity of the ICT test was
     97.2%. CONCLUSION: The ICT assay is a valuable tool for the
     diagnosis of pneumococcal pneumonia, especially for the nonbacteremic
     cases.
CT
     Check Tags: Female; Human; Male
      Adolescence
      Adult
      Aged
      Aged, 80 and over
      Bacteremia: DI, diagnosis
Bacteremia: IM, immunology
      Child
      Child, Preschool
      Middle Age
     *Pneumonia, Pneumococcal: DI, diagnosis
      Pneumonia, Pneumococcal: IM, immunology
     *Polysaccharides, Bacterial: UR, urine
      Predictive Value of Tests
     ANSWER 5 OF 19
L3
                         MEDLINE
                     MEDLINE
ΑN
     2000421114
                PubMed ID: 10878074
DN
     20341579
     Rapid diagnosis of Legionnaires' disease using an
TI
     immunochromatographic assay for Legionella pneumophila serogroup 1
     antigen in urine during an outbreak in the Netherlands. Wever P C; Yzerman E P; Kuijper E J; Speelman P; Dankert J
ΑU
     Department of Medical Microbiology, Academic Medical Center, University
CS
of
     Amsterdam, Amsterdam, The Netherlands.. p.c.wever@amc.uva.nl
     JOURNAL OF CLINICAL MICROBIOLOGY, (2000 Jul) 38 (7) 2738-9.
SO
     Journal code: HSH; 7505564. ISSN: 0095-1137.
CY
     United States
DT
     Journal; Article; (JOURNAL ARTICLE)
LA
     English
FS
     Priority Journals
EM
     200009
     Entered STN: 20000915
ED
     Last Updated on STN: 20000915
     Entered Medline: 20000901
     A new immunochromatographic assay for rapid qualitative
AΒ
     detection of Legionella pneumophila serogroup 1 antigen in urine
specimens
     was used during an outbreak of legionellosis in The Netherlands. The
assay
     seems of the utmost value in providing a rapid diagnosis of Legionnaires'
     disease in patients with severe community-acquired pneumonia in an
     outbreak setting.
CT
     Check Tags: Human
     *Antigens, Bacterial: UR, urine
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Chromatography: MT, methods
     *Disease Outbreaks
     *Immunologic Techniques
      Legionella pneumophila: IM, immunology
     *Legionella pneumophila: IP, isolation & purification
     *Legionnaires' Disease: DI, diagnosis
      Legionnaires' Disease: EP, epidemiology
      Legionnaires' Disease: MI, microbiology
      Netherlands: EP, epidemiology
     ANSWER 6 OF 19
                         MEDIT INF
L3
                    MEDLINE
AN
     2000153062
     20153062
               PubMed ID: 10691203
DN
     Evaluation of a rapid immunochromatographic assay for the
ΤI
     detection of Legionella antigen in urine samples.
     Dominguez J; Gali N; Matas L; Pedroso P; Hernandez A; Padilla E; Ausina V
ΑU
     Servicio de Microbiologia, Hospital Universitari Germans Trias i Pujol,
CS
     Facultad de Medicina de la Universitat Autonoma de Barcelona, Spain..
     jadoming@ns.hugtip.scs.es
     EUROPEAN JOURNAL OF CLINICAL MICROBIOLOGY AND INFECTIOUS DISEASES, (1999
SO
     Dec) 18 (12) 896-8.
     Journal code: EM5; 8804297. ISSN: 0934-9723.
     GERMANY: Germany, Federal Republic of
CY
DT
     Journal; Article; (JOURNAL ARTICLE)
LA
     English
     Priority Journals
FS
     200003
EΜ
     Entered STN: 20000327
ED
     Last Updated on STN: 20000327
     Entered Medline: 20000314
     A new immunochromatographic membrane assay for detecting
AB
     Legionella pneumophila serogroup 1 antigen in urine samples (Binax Now Legionella Urinary Antigen Test; Binax, USA) was evaluated. Its
     sensitivity, specificity and level of agreement with the Binax enzyme
     immunoassay were compared using nonconcentrated and concentrated urine
     samples. The overall agreement between the two tests was 98.1%; the
     specificity of both was 100%. The sensitivity of the
     immunochromatographic assay was 55.5% in nonconcentrated urine and
     97.2% in concentrated urine in comparison with the enzyme immunoassay,
     using concentrated urine as the reference test. This
     immunochromatographic assay screens successfully for Legionella
     pneumophila serogroup 1 soluble antigen in concentrated urine samples.
     Check Tags: Female; Human; Male
CT
      Adult
      Aged
     *Antigens, Bacterial: UR, urine
     *Chromatography: MT, methods
      Evaluation Studies
      Immunoenzyme Techniques
     *Immunologic Techniques
      Legionella pneumophila: IM, immunology
     *Legionella pneumophila: IP, isolation & purification
     *Legionnaires' Disease: DI, diagnosis
      Legionnaires' Disease: MI, microbiology
      Middle Age
      Sensitivity and Specificity
     *Urine: MI, microbiology
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ANSWER 7 OF 19 MEDLINE L3 2000087388 MEDLINE AN PubMed ID: 10618283 DN 20087388 Detection of Francisella tularensis in biological specimens using a TΙ capture enzyme-linked immunosorbent assay, an immunochromatographic handheld assay, and a PCR. Grunow R; Splettstoesser W; McDonald S; Otterbein C; O'Brien T; Morgan C; ΑU Aldrich J; Hofer E; Finke E J; Meyer H Institute of Microbiology, Federal Armed Forces Medical Academy, 80937 CS Munich, Germany.. tb101cn@mail.lrz-muenchen.de CLINICAL AND DIAGNOSTIC LABORATORY IMMUNOLOGY, (2000 Jan) 7 (1) 86-90. SO Journal code: CB7; 9421292. ISSN: 1071-412X. CY United States Journal; Article; (JOURNAL ARTICLE) DT LA English Priority Journals FS EM200003 Entered STN: 20000314 ED Last Updated on STN: 20000314 Entered Medline: 20000302 The early detection of Francisella tularensis, the causative agent of AB tularemia, is important for adequate treatment by antibiotics and the outcome of the disease. Here we describe a new capture enzyme-linked immunosorbent assay (cELISA) based on monoclonal antibodies specific for lipopolysaccharide (LPS) of Francisella tularensis subsp. holarctica and Francisella tularensis subsp. tularensis. No cross-reactivity with Francisella tularensis subsp. novicida, Francisella philomiragia, and a panel of other possibly related bacteria, including Brucella spp., Yersinia spp., Escherichia coli, and Burkholderia spp., was observed. The detection limit of the assay was 10(3) to 10(4) bacteria/ml. This sensitivity was achieved by solubilization of the LPS prior to the In addition, a novel immunochromatographic membrane-based handheld assay (HHA) and a PCR, targeting sequences of the 17-kDa protein (TUL4) gene of F. tularensis, were used in this study. Compared to the cELISA, the sensitivity of the HHA was about 100 times lower and that of the PCR was about 10 times higher. All three techniques were successfully applied to detect F. tularensis in tissue samples of European brown hares (Lepus europaeus). Whereas all infected samples were recognized by the cELISA, those with relatively low bacterial load were partially or not detected by PCR and HHA, probably due to inhibitors or lack of sensitivity. In conclusion, the HHA can be used as a very fast and simple approach to perform field diagnosis to obtain a first hint of an with F. tularensis, especially in emergent situations. In any suspect case, the diagnosis should be confirmed by more sensitive techniques, such as the cELISA and PCR. Check Tags: Comparative Study CTAntibodies, Monoclonal Blotting, Western Brucella: IM, immunology Burkholderia: IM, immunology Cross Reactions \*Enzyme-Linked Immunosorbent Assay Enzyme-Linked Immunosorbent Assay: MT, methods

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Escherichia coli: IM, immunology
      Francisella tularensis: CL, classification
     *Francisella tularensis: GE, genetics
*Francisella tularensis: IP, isolation & purification
     *Immunologic Tests: MT, methods
     *Lipopolysaccharides: IM, immunology
     *Polymerase Chain Reaction
      Polymerase Chain Reaction: MT, methods
      Sensitivity and Specificity
      Yersinia: IM, immunology
     ANSWER 8 OF 19
                          MEDLINE
L3
AN
     1999308407
                      MEDLINE
DN
     99308407
                 PubMed ID: 10383257
     Rapid immunochromatographic assay for diagnosis of tuberculosis:
ΤI
     antibodies detected may not be specific.
CM
     Comment on: J Clin Microbiol. 1998 Nov; 36(11):3443
     Freeman R; Magee J; Barratt A; Wheeler J; Steward M; Lee M; Piggott N
ΑU
     JOURNAL OF CLINICAL MICROBIOLOGY, (1999 Jun) 37 (6) 2111-2.
SO
     Journal code: HSH; 7505564. ISSN: 0095-1137.
CY
     United States
DT
     Commentary
     Letter
LA
     English
     Priority Journals
FS
EM
     199906
     Entered STN: 19990712
ED
     Last Updated on STN: 20000303
     Entered Medline: 19990623
     Check Tags: Human
CT
      Antibodies, Monoclonal
      Antigens, Bacterial: IM, immunology
      Blotting, Western
      Molecular Weight
      Recombinant Proteins: IM, immunology
     *Tuberculosis: DI, diagnosis
Tuberculosis: IM, immunology
Tuberculosis, Pulmonary: DI, diagnosis
Tuberculosis, Pulmonary: IM, immunology
L3
     ANSWER 9 OF 19
                          MEDLINE
ΑN
     1998453872
                     MEDLINE
DN
     98453872
                 PubMed ID: 9780587
ΤI
     Evaluation of immunochromatography-based rapid detection kit for
     fecal Escherichia coli 0157.
     Takeda T; Yamagata K; Yoshida Y; Yoshino K; Nomura T
ΑU
     Department of Infectious Diseases Research, National Children's Medical
CS
     Research Center.
     KANSENSHOGAKU ZASSHI. JOURNAL OF THE JAPANESE ASSOCIATION FOR INFECTIOUS
SO
     DISEASES, (1998 Aug) 72 (8) 834-9.
     Journal code: IJR; 0236671. ISSN: 0387-5911.
CY
     Japan
DT
     Journal; Article; (JOURNAL ARTICLE)
LA
     Japanese
FS
     Priority Journals
     199812
EM
     Entered STN: 19990115
ED
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Last Updated on STN: 19990115 Entered Medline: 19981201

"Quix" is an immunochromatography-based direct detection kit for AB the E. coli 0157 LPS antigen in the patient's stool. The present study

was

conducted to evaluate the efficacy of the kit for rapid diagnosis of enterohemorrhagic Escherichia coli (EHEC) 0157 infection. Sensitivity of the kit was determined using a pure culture of a clinical isolate of E. coli 0157. Analytical sensitivity was found to be  $5 \times 10(5)$  CFU/ml. When compared with the culture method using fecal samples of 64 patients and with bloody diarrhea, sensitivity and specificity were 95.0% (19/20) and 86.4% (38/44), respectively, and overall agreement to culture method was 89.1% (57/64). One patient was found positive by culture method while negative in the present method, where the sample contained a low number

of

is

the cells less than the detection limit. Four of the six patients with a negative result by culture method and positive in the present method,

confirmed E. coli 0157 infection by positive IgM antibody response against

the E. coli O157 LPS. The discrepancy between the two methods seemed to be

attributable to antibiotic administration. In one patient, Salmonella urbana (030(1)30(2)) was detected. The O30(1) antigen of this bacterium

well known to be identical to the E. coli 0157 antigen. When the present method was compared with an ELISA-based E. coli 0157 LPS antigen detection

kit, sensitivity and specificity were 100% (11/11) and 82.1% (23/28), and overall agreement to ELISA method was 87.2% (34/39). From these findings, Quix is useful as a rapid diagnostic kit in the primary clinics, outpatient or bedside use. E. coli 0157 LPS antigen in patient's fecal samples can be detected in about five minutes with this simple procedure. Early diagnosis using such a simple kit will largely contribute for the early treatment and prevention of severe complications of the E. coli

0157

infection.

Check Tags: Female; Human; Male CTAntigens, Bacterial: AN, analysis

Child

Child, Preschool

Escherichia coli Infections: DI, diagnosis

Escherichia coli 0157: IM, immunology

\*Escherichia coli 0157: IP, isolation & purification

Evaluation Studies

\*Feces: MI, microbiology Reagent Kits, Diagnostic

Sensitivity and Specificity

- L3 ANSWER 10 OF 19 MEDLINE
- 1998193869 MEDLINE ΑN
- PubMed ID: 9532690 DN 98193869
- [A comparative study of the information value of the immunofluorescence and immunochromatographic identification of Chlamydia trachomatis antigens in smear material from the cervical canal and of the cytological picture of a vaginal discharge in pregnant women]. Sravnitel'noe izuchenie informativnosti immunofliuorestsentnoi i

Page 20

immunokhromatograficheskoi identifikatsii antigenov Chlamydia trachomatis v soskobnom materiale iz tservikal'nogo kanala i tsitologicheskoi kartiny vaginal'nogo otdeliaemogo u beremennykh.
Zul'karneev R Sh; Kalinin Iu T; Afanas'ev S S; Rubal'skii O V; Denisov L A; Vorob'ev A A; Kiselev V I; Afanas'ev D S; Voronin M V; Afanas'ev M S Health Department of the City Administration of Astrakhan, Russia.

CS Health Department of the City Administration of Astrakhan, Russia.

SO ZHURNAL MIKROBIOLOGII, EPIDEMIOLOGII I IMMUNOBIOLOGII, (1998 Jan-Feb) (1)

64-7.

Journal code: Y90; 0415217. ISSN: 0372-9311.

CY RUSSIA: Russian Federation

DT Journal; Article; (JOURNAL ARTICLE)

LA Russian

ΑU

FS Priority Journals

EM 199806

ED Entered STN: 19980618
Last Updated on STN: 19980618
Entered Medline: 19980608

The cytological picture of the vaginal discharge and scrape material, obtained from 30 pregnant women of the control group (not infected with C.trachomatis) and 61 pregnant women with chlamydiosis; of these, in 42 women the comparative identification of chlamydiae by the methods of direct immunofluorescence and immunochromatography was carried out. Direct immunofluorescence was carried out with the use of a set of reagents "MicroTrak" (USA) and immunochromatographic identification was made with the use of a set of reagents "Clearview Chlamydia" (Britain). The comparison of the results of immunochromatography and direct immunofluorescence revealed that the sensitivity of the set "Clearview Chlamydia" was 100.0% in comparison with the data obtained in the examination of women with the use of the

set

"MicroTrak". The negative results coincided in 90.1% of cases. The common features of the cytological picture of vaginal samples taken from pregnant

women were established. The optimum system of the examination of pregnant women suspected for chlamydiosis, as well as for the evaluation of the effectiveness of its treatment, was proposed.

CT Check Tags: Comparative Study; Female; Human

\*Antigens, Bacterial: AN, analysis
\*Chlamydia Infections: DI, diagnosis
\*Chlamydia trachomatis: IM, immunology
Chromatography
Fluorescent Antibody Technique, Direct
Immunologic Tests
Pregnancy

\*Pregnancy Complications, Infectious: DI, diagnosis Sensitivity and Specificity

Time Factors

\*Vaginal Discharge: DI, diagnosis

\*Vaginal Smears: MT, methods

L3 ANSWER 11 OF 19 MEDLINE

AN 1998134302 MEDLINE

DN 98134302 PubMed ID: 9474027

TI Development of FlexSure HP--an immunochromatographic method to detect antibodies against Helicobacter pylori.

AU Schrier W H; Schoengold R J; Baker J T; Norell J L; Jaseph C L; Okin Y; Doe J Y; Chandler H

Page 21

SmithKline Diagnostics, Inc., Palo Alto, CA 94303, USA. CS CLINICAL CHEMISTRY, (1998 Feb) 44 (2) 293-8. SO Journal code: DBZ; 9421549. ISSN: 0009-9147. CY United States Journal; Article; (JOURNAL ARTICLE) DTLA English FS Priority Journals 199803 EMEntered STN: 19980326 ED Last Updated on STN: 19980326 Entered Medline: 19980316 We describe a solid-phase immunochromatographic serologic test, AB FlexSure HP, to detect IgG antibodies against Helicobacter pylori. H. pylori colonize the stomach and proximal duodenum, cause ulcer disease and mucosa-associated lymphoid tissue lymphoma, and have a role in the development of other disorders, including gastric adenocarcinoma. FlexSure HP consists of a test strip, conjugate pad, and absorbent pad, in a novel reverse-flow chromatography format. In these studies, FlexSure HP was demonstrated to be specific for IgG antibodies against H. pylori. The reactive cutoff of the test was consistent with [13C]urea breath test and commercially available ELISAs. FlexSure HP had 94% sensitivity, 88% specificity, and 91% accuracy relative to [13C]urea breath test; and 95% sensitivity, 94% specificity, and 95% overall agreement relative to high-molecular-mass cell-associated protein enzyme immunoassay (HM-CAP EIA). FlexSure HP is a simple-to-perform, visually read test requiring no specialized training, equipment, or instrumentation, and yields rapid, accurate, qualitative results. Check Tags: Female; Human; Male CTAdult \*Antibodies: BL, blood Antigens, Bacterial: BL, blood Antigens, Bacterial: IM, immunology Breath Tests Cross Reactions: IM, immunology \*Helicobacter Infections: BL, blood Helicobacter Infections: IM, immunology \*Helicobacter pylori: IM, immunology Immunosorbents: ME, metabolism Middle Age ROC Curve Reagent Kits, Diagnostic Reproducibility of Results Sensitivity and Specificity Stomach: MI, microbiology Urea: AN, analysis ANSWER 12 OF 19 L3 MEDLINE 96388854 MEDLINE ANPubMed ID: 8796254 96388854 DN Clinical evaluation of a rapid immunochromatographic assay based TΙ on the 38 kDa antigen of Mycobacterium tuberculosis on patients with pulmonary tuberculosis in China. Comment in: Tuber Lung Dis. 1997;78(1):85 CM Cole R A; Lu H M; Shi Y Z; Wang J; De-Hua T; Zhou A T ΑU ICT Diagnostics, Sydney, Australia. CS

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TUBERCLE AND LUNG DISEASE, (1996 Aug) 77 (4) 363-8.
SO
     Journal code: A8C; 9212467. ISSN: 0962-8479.
     SCOTLAND: United Kingdom
CY
     Journal; Article; (JOURNAL ARTICLE)
DT
LA
     English
FS
     Priority Journals
EM
     199610
ED
     Entered STN: 19961106
     Last Updated on STN: 19990129
     Entered Medline: 19961022
     SETTING: A rapid membrane-based antibody assay capable of diagnosing
AB
     pulmonary tuberculosis within 15 min has been developed using the 38~\rm kDa antigen from Mycobacterium tuberculosis. OBJECTIVE: To determine the
     specificity and sensitivity of this assay and evaluate its usefulness in
а
     clinical setting. DESIGN: Sera from patients with active pulmonary
     tuberculosis were obtained from three hospitals in China. The control
     groups consisted of patients who were diagnosed with lung diseases other
     than tuberculosis and healthy subjects. RESULTS: Antibody was detected in
     54 of 61 (89%) sputum positive patients and 67 of 91 (74%) sputum
negative
     patients who had been clinically diagnosed as having active pulmonary
     tuberculosis. Five out of 56 (9%) patients with respiratory diseases
other
     than tuberculosis and 1 out of 30 (3%) healthy controls had a positive
     antibody response. The overall specificity of the assay was 93% and the
     positive predictive value was 95%. We conclude that this assay is rapid,
     sensitive and specific and will be a valuable aid in the clinical
     diagnosis of pulmonary tuberculosis.
     Check Tags: Female; Human; Male
CT
     *Antigens, Bacterial: AN, analysis
     *Immunoassay: MT, methods
      Middle Age
     *Mycobacterium tuberculosis: IM, immunology
      Mycobacterium tuberculosis: IP, isolation & purification
      Predictive Value of Tests
      Respiratory Tract Diseases: IM, immunology
      Sensitivity and Specificity Sputum: MI, microbiology
      Time Factors
     *Tuberculosis, Pulmonary: DI, diagnosis
      Tuberculosis, Pulmonary: MI, microbiology
     ANSWER 13 OF 19
                          MEDLINE
L3
                  MEDLINE
     94154898
AN
                 PubMed ID: 8111517
     94154898
DN
ΤI
     Comparison of an immunochromatographic method for rapid
     identification of group A streptococcal antigen with culture method.
     Ehrlich T P; Schwartz R H; Wientzen R; Thorne M M
ΑU
     Department of Family Practice, Medical College of Virginia, Fairfax
CS
Family
     Practice Center Inc.
     ARCHIVES OF FAMILY MEDICINE, (1993 Aug) 2 (8) 866-9.
SO
     Journal code: BX6; 9300357. ISSN: 1063-3987.
     United States
CY
     Journal; Article; (JOURNAL ARTICLE)
DT
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LA English

FS Priority Journals

EM 199403

ED Entered STN: 19940406

Last Updated on STN: 19940406 Entered Medline: 19940330

AB OBJECTIVES: To compare the sensitivity and specificity of Concise Strep A (Hybritech, San Diego, Calif), an immunochromatographic group A streptococcal rapid antigen detection system, with a two-plate culture method for the diagnosis of streptococcal pharyngitis, and to evaluate

the

need for routine back-up culture when this rapid test is used. DESIGN: Throat cultures were obtained from 351 children with acute pharyngitis by duplicate rayon-tipped swabs held in parallel and vigorously rubbed against both tonsils and the posterior pharyngeal wall. One swab was tested for group A streptococcal antigen by a registered licensed laboratory technologist in the pediatrician's office. The other swab was streaked over each of two sheep blood agar plates, one of which was enhanced with trimethoprim in combination with sulfamethoxazole. The

plain

sheep blood agar plate was then incubated in a candle-extinguish jar. The enhanced agar plate was placed in a gas-pack anaerobic jar. Both plates were incubated for up to 48 hours at 35 degrees C. SETTING: A six-person group pediatric practice. PARTICIPANTS: Three hundred fifty-one children. RESULTS: The Concise Strep A antigen detection test produced 129 positive results. Only six of the 129 were not confirmed by culture method. There were four false-negative rapid streptococcal antigen detection test results, all of which were found after a single overnight incubation. The sensitivity for the Concise Strep A test was 96.9% and the specificity

was

97.4%. The plain 5% sheep blood agar plate (without trimethoprim and sulfamethoxazole), which was incubated in a candle-extinguish jar, identified 123 (97%) of the 127 positive throat cultures. The second 24-hour incubation and use of trimethoprim and sulfamethoxazole agar were not rewarding for this study. CONCLUSIONS: Concise Strep A, a polyclonal antibody test, in conjunction with a color immunochromatographic assay for soluble streptococcal carbohydrate antigen A appears to be accurate, sensitive, and specific when throat swabs are carefully

obtained

and when qualified, licensed laboratory technologists perform the procedure. Further studies should be done to confirm our findings, especially when nurses or office staff perform the rapid test procedure

in

the office setting. If our findings are confirmed, the use of back-up cultures for negative rapid test results obtained using Concise Strep A would be unnecessary.

CT Check Tags: Comparative Study; Human; Support, Non-U.S. Gov't Acute Disease

\*Antigens, Bacterial: AN, analysis

Bacteriological Techniques

Child

Chromatography

False Negative Reactions

Immunoassay

\*Pharyngitis: DI, diagnosis Pharyngitis: MI, microbiology Pharynx: MI, microbiology

Page 24

Reagent Kits, Diagnostic Sensitivity and Specificity \*Streptococcal Infections: DI, diagnosis Streptococcus pyogenes: IM, immunology \*Streptococcus pyogenes: IP, isolation & purification ANSWER 14 OF 19 MEDLINE L3 . MEDLINE ΑN 93036353 DN 93036353 PubMed ID: 1416045 Kinetic chromatographic sequential addition immunoassays using protein A TI affinity chromatography. ΑU Cassidy S A; Janis L J; Regnier F E Department of Chemistry, Purdue University, West Lafayette, Indiana CS 47907. NC GM 25431 (NIGMS) ANALYTICAL CHEMISTRY, (1992 Sep 1) 64 (17) 1973-7. SO Journal code: 4NR; 0370536. ISSN: 0003-2700. CYUnited States Journal; Article; (JOURNAL ARTICLE) DT LA English Priority Journals FSEM199210 Entered STN: 19930122 ΕD Last Updated on STN: 19930122 Entered Medline: 19921026 A new type of chromatographic immunoassay based on sequential addition is AΒ described. On a protein A column, the antibody, the sample containing the antigen, and then a known amount of antigen are sequentially injected. This assay is designed to shorten analysis times and reduce complexity of dual-column chromatographic immunoassays, circumvent desorption buffer interferences common to affinity chromatography, and eliminate the need for tagged molecules. This new technique is named kinetic immunochromatography sequential addition (KICQA). Because of its kinetic nature, flow rate will have a large effect on KICQA, and the impact of changing flow rate is studied extensively. By use of various amounts of antibody, the dynamic range of KICQA is shown to be selectable over 2.5 orders of magnitude. Finally, KICQA was used to determine transferrin and albumin in human serum. Both analytes show good agreement with their respective reference methods, and an albumin assay was performed in under 1 min. Check Tags: Support, U.S. Gov't, P.H.S. CTAntigen-Antibody Complex \*Apoproteins: AN, analysis \*Chromatography, Affinity \*Immunoassay \*Serum Albumin: AN, analysis \*Staphylococcal Protein A: CH, chemistry \*Transferrin: AN, analysis ANSWER 15 OF 19 MEDLINE L3 AN90028644 MEDLINE PubMed ID: 2679905 DN 90028644 [Monoclonal antibodies cross-reacting with fibroblasts of interstitial ΤI connective tissue of the myocardium and cell wall protein antigens of group A Streptococcus]. Monoklonal'nye antitela, perekrestno reagiruiushchie s fibroblastami interstitsial'noi soedinitel'noi tkani miokarda i belkovymi antigenami Page 25

kletochnoi stenki streptokokka gruppy A. Abyzov V N; Drobyshevskaia E I; Liampert I M; Borodiiuk N A; Panasiuk A F ΑU BIULLETEN EKSPERIMENTALNOI BIOLOGII I MEDITSINY, (1989 Jul) 108 (7) 74-6. SO Journal code: A74; 0370627. ISSN: 0365-9615. CY Journal; Article; (JOURNAL ARTICLE) DTLA Russian Priority Journals FS EM198912 Entered STN: 19900328 ED Last Updated on STN: 19900328 Entered Medline: 19891201 Monoclonal antibodies (MCA) B6/5 and C5/3 were obtained after AΒ immunization of BALB/c mice with the protein non-type-specific antigens (NTSA) of streptococcal group A cell wall. MCA B6/5 in the indirect immunofluorescence react with human and animal interstitial connective tissue (ICT) of the myocardium and human fibroblast culture cells. MCA C5/3 react with the bands of muscle fibers of the myocardium. MCA B6/5 and C5/3 are autoantibodies. It was revealed that these MCA are directed to two streptococcal cross-reacting antigens (CRA). Production of B6/5 and C5/3, apparently, does not depend on the possibility of some streptococcal antigens to bind fibrinogen. Bound immunoglobulins were not revealed in the ICT and in the muscle fibres by the cultivation of the C5/3 monoclone. Firstly it was stated that, MCA B6/5, reacting with fibroblasts and with streptococcal CRA, are capable to fix in the ICT of myocardium, what is typical for the phenomenon described in rheumatic fever. CTCheck Tags: Animal; Human \*Antibodies, Monoclonal: IM, immunology \*Antigens, Bacterial: IM, immunology \*Autoantibodies: AN, analysis Cross Reactions Fibroblasts: IM, immunology Fluorescent Antibody Technique Mice Mice, Inbred BALB C \*Myocardium: IM, immunology Rheumatic Fever: IM, immunology \*Streptococcus pyogenes: IM, immunology L3 ANSWER 16 OF 19 MEDLINE ΑN 81035609 MEDLINE PubMed ID: 6158813 DN 81035609 [T-proteins of Streptococcus pyogenes. II. Communication: Preparation of ΤI specific immunoadsorbents for isolation of anti-T-antibodies and investigations of the T4/24...pattern (author's transl)]. T-Proteine des Streptococcus pyogenes. II. Mitteilung: Praparation spezifischer Immunadsorbentien zur Isolierung von T-Antikorpern und Untersuchungen de T4/24...Agglutinationsgruppe. Schmidt K H; Kuhnemund O; Kohler W ΑU ZENTRALBLATT FÜR BAKTERIOLOGIE. 1. ABT. ORIGINALE. A: MEDIZINISCHE SO MIKROBIOLOGIE, INFEKTIONSKRANKHEITEN UND PARASITIOLOGIE, (1980) 246 (4) 489-98.

Journal code: Y5P; 8005748. ISSN: 0172-5599.

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GERMANY, WEST: Germany, Federal Republic of
CY
    Journal; Article; (JOURNAL ARTICLE)
DT
LA
    German
     Priority Journals
FS
EM
     198012
     Entered STN: 19900316
ED
     Last Updated on STN: 19900316
     Entered Medline: 19801218
    The T4-antigen(s) in Streptococcus pyogenes standard type strains of the
ΑB
     T4/24...pattern was (were) investigated in regard to the uniformity or
    diversity of the antigen(s) in this T-complex. Tryptic T-extracts of
types
     4, 24, 26, 28, 29, 46, 48, 60 were purified by ion exchange
chromatography
    on DEAE-cellulose. Anti-T4-antibodies were isolated by immunoadsorption
     chromatography on AH-sepharose linked T4-antigen. Purified T4-antigen
     showed in SDS-electrophoresis a similar multiple molecular size structure
     as T1-antigen described earlier. Comparative serological studies of
     T-antigens of types 4, 24, 29 and 46 revealed reactions of identity to
     anti-T4-antibodies in Ouchterlony tests. Extracts of types 26, 28, 48 and
     60 did not precipitate with anti-T4-antibodies, but types 28 and 48
showed
    crossreaction to the relevant antisera (anti-T28 and anti-T48, resp.)
    obviously caused by traces of R-28 antigen in both antigen preparations.
     Strains of the types 4, 24, 29, 46, 48 and 60 were agglutinated by
     anti-T4-antibodies. The reaction could be inhibited by T4-antigen. The
     strains of type 26 and 28 used in our experiments did not contain
    T4-antigen. Agglutination as well as immunoprecipitation reactions with
     specific antibodies prepared by immunochromatography proved the
     existence of common T4-antigenic determinants in type 4, 24, 29, 46, 48
     and 60.
CT
    Check Tags: Animal
     Antibodies, Bacterial: IM, immunology
     *Antibodies, Bacterial: IP, isolation & purification
     *Antigen-Antibody Complex
     *Antigens, Bacterial: IM, immunology
     Antigens, Bacterial: IP, isolation & purification
     *Bacterial Proteins: IM, immunology
      Epitopes
      Immunodiffusion
     *Immunosorbents
      Rabbits
     *Streptococcus pyogenes: IM, immunology
     ANSWER 17 OF 19
                         MEDLINE
L3
                  MEDLINE
AN
     80243108
                PubMed ID: 6994834
DN
     80243108
     [Study of cross reactions between group A streptococcal antigens and
TΙ
     cardiac interstitial connective tissue fibroblasts of different species
of
    mammals].
     Izuchenie perekrestnykh reaktsii mezhdu antigenami streptokokka gruppy A
i
     fibroblastami interstitsial'noi soedinitel'noi tkani serdtsa
    mlekopitaiushchikh raznykh vidov.
     Kochetkova E V; Liampert I M; Kolesnikova V Iu; Semenova E N
ΑU
     BIULLETEN EKSPERIMENTALNOI BIOLOGII I MEDITSINY, (1980 May) 89 (5) 582-4.
                                                                        Page 27
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Journal code: A74; 0370627. ISSN: 0365-9615.
CY
     USSR
     Journal; Article; (JOURNAL ARTICLE)
DT
LA
     Russian
     Priority Journals
FS
EM
     198010
     Entered STN: 19900315
ED
     Last Updated on STN: 19900315
     Entered Medline: 19801027
    It has been revealed by indirect immunofluorescence that sera of rabbits
     immunized with group A streptococcal non-type-specific antigens and F
     (ab')2 fragments of IgG obtained from these sera intensively react with
     fibroblasts of interstitial connective tissue (ICT) of the
     myocardium of humans and different mammalian species with the exception
οf
     rabbits. Negative or weak reactions with the same sera and F (ab')2
     fragments were observed with fibroblasts of ICT of nonimmunized
     rabbits. Myocardial ICT of immunized animals showed bound
     immunoglobulins. This model can be used for making clear the reason for
     presence in sera of rheumatic patients circulating antibodies only to
     heterologous but not to homologous ICT of the myocardium
     regardless of the revealing bound immunoglobulins in the ICT of
     these patients.
     Check Tags: Animal; Comparative Study; Human
CT
      Antigens, Bacterial: IM, immunology
      Autoantibodies: AN, analysis
      Cattle
     *Cross Reactions
      Fibroblasts: IM, immunology
      Fluorescent Antibody Technique
      Guinea Pigs
      Heart: EM, embryology
      Immunoglobulins, Fab: IM, immunology
     Myocardium: CY, cytology *Myocardium: IM, immunology
      Rabbits: IM, immunology
      Rheumatic Fever: IM, immunology
      Species Specificity
     *Streptococcus pyogenes: IM, immunology
      Swine: IM, immunology
L3
     ANSWER 18 OF 19
                         MEDLINE
                  MEDLINE
AN.
     80181173
                PubMed ID: 6989752
     80181173
DN
     Mitogenic and antigenic properties of group A streptococcal M protein
TI
     preparations.
     Knoll H; Kuhnemund O; Havlicek J
AU
     IMMUNOBIOLOGY, (1980 Jan) 156 (4-5) 537-48.
SO
     Journal code: GH3; 8002742. ISSN: 0171-2985.
     GERMANY, WEST: Germany, Federal Republic of
CY
\mathsf{DT}
     Journal; Article; (JOURNAL ARTICLE)
LA
     English
     Priority Journals
FS
     198007
EΜ
     Entered STN: 19900315
ED
     Last Updated on STN: 19900315
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Page 28

Entered Medline: 19800728 AB The separation of a mitogenic substance in M protein preparations of Streptococcus pyogenes, type 1 and type 12, is described. The isolation was achieved by gel chromatography on Biogel A 0.5 m, and by immunochromatography on immobilized type specific antibodies. In the delayed cutaneous hypersensitivity test the immunochromatography purified M 1 protein caused erythema but no unspecific mitogenicity could be found by lymphocyte transformation test. In neutralization experiments the mitogenic activity of M protein preparations was specifically inhibited by anti-erythrogenic toxin antisera. Check Tags: Animal; Human Adult Antibody Specificity \*Antigens, Bacterial \*Bacterial Proteins: IM, immunology Bacterial Proteins: IP, isolation & purification Chromatography, Gel Fractionation Hydrochloric Acid: IM, immunology Hypersensitivity, Delayed: IM, immunology Middle Age \*Mitogens: PD, pharmacology Neutralization Tests Precipitation Rabbits \*Streptococcus pyogenes: IM, immunology ANSWER 19 OF 19 MEDLINE L3MEDLINE AN80061440 PubMed ID: 388938 80061440 DN Immunoelectron microscopic localization of T proteins in the cell wall of TΤ Streptococcus pyogenes. ΑU Wagner B; Schmidt K H; Wagner M ZENTRALBLATT FUR BAKTERIOLOGIE, PARASITENKUNDE, INFEKTIONSKRANKHEITEN UND S0HYGIENE. ERSTE ABTEILUNG ORIGINALE. REIHE A: MEDIZINISCHE MIKROBIOLOGIE UND PARASITOLOGIE, (1979 Jul) 244 (2-3) 192-201. Journal code: Y52; 0331570. ISSN: 0300-9688. GERMANY, WEST: Germany, Federal Republic of CY DT Journal; Article; (JOURNAL ARTICLE) LA English FS Priority Journals EΜ 198001 Entered STN: 19900315 Last Updated on STN: 19900315 Entered Medline: 19800128 T proteins of Streptococcus pyogenes (group A streptococci) were AΒ localized by means of immunochromatographical isolated anti-T-antibodies. For the electron microscopical detection both the direct and the indirect immunoferritin techniques were used. The arrangement of the ferritin particles showed, that the T proteins are evenly distributed on the whole cell surface. They are immediately bound to the outer layer of the cell wall or to only short filaments. On isolated cell walls the T protein was detected only on the outer surface. CTAgglutination Tests

Antigens, Bacterial

\*Bacterial Proteins
\*Cell Wall
\*Streptococcus pyogenes: UL, ultrastructure